Express Mail" mailing label number EM 062 546 604 US. I hereby certify that this document and referenced attachments are being deposited with the United States Postal Service "Express Mail Post Office to Addressee" service under CFR § 1.10, addressed to: Assistant Commissioner for Patents, ox Sequence, Washington, D.C. 20231 on December 31, 1997.

Emma Durrell

Assistant Commissioner for Patents **Box Sequence** Washington, D.C. 20231

Transmitted herewith for filing is the patent application of:

Inventors: Preeti Lal, Jennifer L. Hillman, Neil C. Corley, Karl J. Guegler, Mariah Baugh, Susan Sather and Purvi Shah

Title:

### HUMAN SIGNAL PEPTIDE-CONTAINING PROTEINS

Enclosed are:

Return postcard, XXXXXXXXX

- 117 Pages of Specification;
- 169 Pages of Sequence Listing,
- Pages of Claims,
- Page of Abstract;
- -0- Pages of Figures
  - 5 Pages Unexecuted Declaration and Power of Attorney; and
  - 1 Page of Sequence Listing Statement and one (1) Computer-Readable Diskette.

Fee Calculation - The fee has been calculated as follows:

### CLAIMS AS FILED (fees computed under § 1.9(f))

Claims	Number Filed	Minus	Number Extra	Other I Small E Rate	
Total Claims	23	-20	3	x \$22	\$ 66 00
Indep. Claims	2	-3	-0-	x \$82	\$ -0-
Multiple Depe	ndent Claim(s), if any		+ \$270	· · · · · · · · · · · · · · · · · · ·	\$ -0-

TOTAL FILING FEE

**\$ 856.00** 

The Commissioner is hereby authorized to charge Deposit Account No. 09-0108 in the amount of § 856.00

The Commissioner is hereby authorized to charge any additional fees required under 37 C.F.R. § 1.16 and 1.17, or credit any overpayment to Deposit Account No. 09-0108. A duplicate of this sheet is enclosed.

Respectfully submitted,

INCYTE PHARMACEUTICALS, INC.

Reg. No. 41,201

3174 Porter Drive

Palo Alto, California 94304 Phone: (650) 855-0555

Fax: (650) 845-4166



The first thing the first the first that the first that the first the first that the first that

"Express Mail" mailing label number EM 062 546 604 US. I hereby certify that this document and referenced attachments are being deposited with the United States Postal Service "Express Mail Post Office to Addressee" service under 37 CFR § 1.10, addressed to: Assistant Commissioner for Patents, Box Sequence, Washington, D.C. 20231 on December 31, 1997.

By: Lynna Durell
Printed: Emma Durrell

### IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:

Lal et al.

Title:

**HUMAN SIGNAL PEPTIDE-CONTAINING PROTEINS** 

Serial No.:

To Be Assigned

Filing Date:

Herewith

Examiner:

To Be Assigned

Group Art Unit: To Be Assigned

**Assistant Commissioner for Patents** 

**Box Sequence** 

Washington, D.C. 20231

### SUBMISSION UNDER 37 CFR § 1.821-1.825 SEQUENCE LISTING

Sir:

In accordance with the requirements of 37 CFR § 1.821-1.825, Applicants hereby submit one diskette containing the computer-readable information for the Sequence Listing of the above-identified application. The diskette complies with the requirements of 37 CFR § 1.824 and is IBM PC compatible using a PC-DOS/MS-DOS 6.2 operating system with Fastseq software version 2.0.

Contained within the application, as filed, just before the claim section, is the Sequence Listing paper copy of the sequences disclosed in the application.

The content of the Sequence Listing paper copy is identical to the computer-readable copy, as required under 37 CFR § 1.821(f).

Respectfully submitted,

INCYTE PHARMACEUTICALS, INC.

Date: December 31,199

Sheela Mohan-Peterson

Reg. No. 41,201

3174 Porter Drive

Palo Alto, California 94304 Phone: (650) 855-0555

Fax: (650) 845-4166

25

5

### HUMAN SIGNAL PEPTIDE-CONTAINING PROTEINS

### FIELD OF THE INVENTION

This invention relates to nucleic acid and amino acid sequences of human signal peptide-containing proteins and to the use of these sequences in the diagnosis, treatment, and prevention of cancer and immunological disorders.

### **BACKGROUND OF THE INVENTION**

Protein transport is an essential process for all living cells. Transport of an individual protein usually occurs via an amino-terminal signal sequence which directs, or targets, the protein from its ribosomal assembly site to a particular cellular or extracellular location. Transport may involve any combination of several of the following steps: contact with a chaperone, unfolding, interaction with a receptor and/or a pore complex, addition of energy, and refolding. Moreover, an extracellular protein may be produced as an inactive precursor. Once the precursor has been exported, removal of the signal sequence by a signal peptidase and posttranslational processing (e.g., glycosylation or phosphorylation) activates the protein. Signal sequences are common to receptors, matrix molecules (e.g., adhesion, cadherin, extracellular matrix, integrin, and selectin), cytokines, hormones, growth and differentiation factors, neuropeptides, vasomediators, phosphokinases, phosphotipases, phospholipases, phosphodiesterases, G and Ras-related proteins, ion channels, transporters/pumps, proteases, and transcription factors.

G-protein coupled receptors (GPCRs) are a superfamily of integral membrane proteins which transduce extracellular signals. GPCRs include receptors for biogenic amines, e.g., dopamine, epinephrine, histamine, glutamate (metabotropic effect), acetylcholine (muscarinic effect), and serotonin; for lipid mediators of inflammation such as prostaglandins, platelet activating factor, and leukotrienes; for peptide hormones such as calcitonin, C5a anaphylatoxin, follicle stimulating hormone, gonadotropin releasing hormone, neurokinin, oxytocin, and thrombin; and for sensory signal mediators, e.g., retinal photopigments and olfactory stimulatory molecules.

25

30

### PF-0459 US

The structure of these highly-conserved receptors consists of seven hydrophobic transmembrane regions, cysteine disulfide bridges between the second and third extracellular loops, an extracellular N-terminus, and a cytoplasmic C-terminus. Three extracellular loops alternate with three intracellular loops to link the seven transmembrane regions. The N-terminus interacts with ligands, the disulfide bridge interacts with agonists and antagonists, and the large third intracellular loop interacts with G proteins to activate second messengers such as cyclic AMP (cAMP), phospholipase C, inositol triphosphate, or ion channel proteins. The most conserved parts of these proteins are the transmembrane regions and the first two cytoplasmic loops. A conserved, acidic-Arg-aromatic triplet present in the second cytoplasmic loop may interact with the G proteins. The consensus pattern, [GSTALIVMYWC]-[GSTANCPDE]-{EDPKRH}-x(2)-[LIVMNQGA]-x(2)-[LIVMFT]-[GS TANC]-[LIVMFYWSTAC]-[DENH]- R-[FYWCSH]-x(2)-[LIVMNQGA]-x(2)-[LIVMFT]-[GS TANC]-[LIVMFYWSTAC]-[DENH]- R-[FYWCSH]-x(2)-[LIVM] is characteristic of most proteins belonging to this superfamily. (Watson, S. and Arkinstall, S. (1994) The G-protein Linked Receptor Facts Book, Academic Press, San Diego, CA, pp. 2-6; and Bolander, F.F. (1994) Molecular Endocrinology, Academic Press, San Diego, CA, pp. 8-19.)

Tetraspanins are a superfamily of membrane proteins which facilitate the formation and stability of cell-surface signaling complexes containing lineage-specific proteins, integrins, and other tetraspanins. They are involved in cell activation, proliferation (including cancer), differentiation, adhesion, and motility. These proteins cross the membrane four times, have conserved intracellular N- and C-termini and an extracellular, non-conserved hydrophilic domain. Three highly conserved polar amino acids are located in the transmembrane domains (TM), an asparagine in TM1 and a glutamate or glutamine in TM3 and TM4. Two to three conserved charged residues, including a glutamic acid residue, are present in the cytoplasmic loop between TM2 and TM3. The extracellular loop between TM3 and TM4 contains four conserved cysteine residues: two in a conserved CCG motif located about 50 residues C-terminal to TM3; one, often preceded by glycine, 11 residues N-terminal to TM4; and one in the extracellular loop may be found in a PXSC motif.

Tetraspanins include, e.g., platelet and endothelial cell membrane proteins, leukocyte surface proteins, tissue specific and tumorous antigens, and the retinitis pigmentosa-associated gene peripherin. (Maecker, H.T. et al. (1997) FASEB J. 11:428-442.) Matrix proteins (Mps)

25

30

5

### PF-0459 US

function in formation, growth, remodeling and maintenance of tissues and as important mediators and regulators of the inflammatory response. The expression and balance of MPs may be perturbed by biochemical changes that result from congenital, epigenetic, or infectious diseases. In addition, MPs affect leukocyte migration, proliferation, differentiation, and activation in immune response.

MPs encompass a variety of proteins and their functions. Extracellular matrix (ECM) proteins are multidomain proteins that play an important role in the diverse functions of the ECM. ECM proteins are frequently characterized by the presence of one or more domains which may include collagen-like domains, EGF-like domains, immunoglobulin-like domains, fibronectin-like domains, vWFA-like modules. (Ayad, S. et al. (1994) The Extracellular Matrix Facts Book, Academic Press, San Diego, CA, pp. 2-16.) Cell adhesion molecules (CAMs) have been shown to stimulate axonal growth through homophilic and/or heterophilic interactions with other molecules. In addition, interactions between adhesion molecules and their receptors can potentiate the effects of growth factors upon cell biochemistry via shared signaling pathways. (Ruoslahti, E. (1997) Kidney Int. 51:1413-1417.) Cadherins comprise a family of calcium-dependant glycoproteins that function in mediating cell-cell adhesion in solid tissues of multicellular organisms. Integrins are ubiquitous transmembrane adhesion molecules that link cells to the ECM by interacting with the cytoskeleton. Integrins also function as signal transduction receptors and stimulate changes in intracellular calcium levels and protein kinase activity. (Sjaastad, M.D. and Nelson, W.J. (1997) BioEssays 19:47-55.)

Lectins are proteins characterized by their ability to bind carbohydrates on cell membranes by means of discrete, modular carbohydrate recognition domains, CRDs. (Kishore, U. et al. (1997) Matrix Biol. 15:583-592.) Certain cytokines and membrane-spanning proteins have CRDs which may enhance interactions with extracellular or intracellular ligands, with proteins in secretory pathways, or with molecules in signal transduction pathways. The lipocalin superfamily constitutes a phylogenetically conserved group of more than forty proteins that function by binding to and transporting a variety of physiologically important ligands. Members of this family function as carriers of retinoids, odorants, chromophores, pheromones, and sterols, and a subset of these proteins may be multifunctional, serving as either a biosynthetic enzyme or as a specific enzyme inhibitor.

25

30

5

(Tanaka, T. et al. (1997) J. Biol. Chem. 272:15789-15795; and van't Hof, W. et al. (1997) J. Biol. Chem. 272:1837-1841.) Selectins are a family of calcium ion-dependent lectins expressed on inflamed vascular endothelium and the surface of some leukocytes. They mediate rolling movement and adhesive contacts between blood cells and blood vessel walls. The structure of the selectins and their ligands supports the type of bond formation and

dissociation that allows a cell to roll under conditions of flow. (Rossiter, H. et al. (1997) Mol.

Med. Today 3:214-222.)

Protein kinases regulate many different cell proliferation, differentiation, and signaling processes by adding phosphate groups to proteins. Reversible protein phosphorylation is a key strategy for controlling protein functional activity in eukaryotic cells. The high energy phosphate which drives this activation is generally transferred from adenosine triphosphate molecules (ATP) to a particular protein by protein kinases and removed from that protein by protein phosphatases. Phosphorylation occurs in response to extracellular signals, cell cycle checkpoints, and environmental or nutritional stresses. Protein kinases may be roughly divided into two groups; protein tyrosine kinases (PTKs) which phosphorylate tyrosine residues, and serine/threonine kinases (STKs) which phosphorylate serine or threonine residues. A few protein kinases have dual specificity. A majority of kinases contain a similar 250-300 amino acid catalytic domain which can be further divided into eleven subdomains. The N-terminal domain, which contains subdomains I to IV, generally folds into a two-lobed structure which binds and orients the ATP (or GTP) donor molecule. The larger C terminal domain, which contains subdomains VIA to XI, binds the protein substrate and carries out the transfer of the gamma phosphate from ATP to the hydroxyl group of the target amino acid residue. Subdomain V links the two domains. Each of the 11 subdomains contain specific residues and motifs that are characteristic and are highly conserved. (Hardie, G. and Hanks, S. (1995) The Protein Kinase Facts Book, Vol I, pp. 7-47, Academic Press, San Diego, CA.)

Protein phosphatases remove phosphate groups from molecules previously modified by protein kinases thus participating in cell signaling, proliferation, differentiation, contacts, and oncogenesis. Protein phosphorylation is a key strategy used to control protein functional activity in eukaryotic cells. The high energy phosphate is transferred from ATP to a protein

30

5

10

by protein kinases and removed by protein phosphatases. There appear to be three, evolutionarily-distinct protein phosphatase gene families: protein phosphatases (PPs); protein tyrosine phosphatases (PTPs); and acid/alkaline phosphatases (APs). PPs dephosphorylate phosphoserine/threonine residues and are an important regulator of many cAMP mediated, hormone responses in cells. PTPs reverse the effects of protein tyrosine kinases and therefore play a significant role in cell cycle and cell signaling processes. Although APs dephosphorylate substrates <u>in vitro</u>, their role <u>in vivo</u> is not well known. (Carbonneau, H. and Tonks, N.K. (1992) Annu. Rev. Cell Biol. 8:463-493.)

Protein phosphatase inhibitors control the activities of specific phosphatases. A specific inhibitor of PP-I, I-1, has been identified that when phosphorylated by cAMP-dependent protein kinase (PKA) specifically binds to PP-I and inhibits its activity. Since PP-I is dephosphoryles many of the proteins phosphorylated by PKA, activation of I-1 by PKA serves to amplify the effects of PKA and the many cAMP-dependent responses mediated by PKA. In addition, since PP-I also dephosphorylates many phosphoryoteins that are not phosphorylated by PKA, I-1 activation serves to exert cAMP control over other protein phosphorylations. I<sub>1</sub>PP2A is a specific and potent inhibitor of PP-IIA. (Li, M. et al. (1996) Biochemistry 35:6998-7002.) Since PP-IIA is the main phosphatase responsible for reversing the phosphorylations of serine/threonine kinases, I<sub>1</sub>PP2A has broad effects in controlling protein phosphorylations.

Cyclic nucleotides (cAMP and cGMP) function as intracellular second messengers to transduce a variety of extracellular signals, including hormones, and light and neurotransmitters. Cyclic nucleotide phosphodiesterases (PDEs) degrade cyclic nucleotides to their corresponding monophosphates, thereby regulating the intracellular concentrations of cyclic nucleotides and their effects on signal transduction. At least seven families of mammalian PDEs have been identified based on substrate specificity and affinity, sensitivity to cofactors and sensitivity to inhibitory drugs. (Beavo, J.A. (1995) Physiological Reviews 75: 725-748.) PDEs are composed of a catalytic domain of ~270 amino acids, an N-terminal regulatory domain responsible for binding cofactors and, in some cases, a C-terminal domain with unknown function. Within the catalytic domain, there is approximately 30% amino acid identity between PDE families and ~85-95% identity between isozymes of the same family.

30

5

10

### PF-0459 US

Furthermore, within a family there is extensive similarity (>60%) outside the catalytic domain, while across families there is little or no sequence similarity. A variety of diseases have been attributed to increased PDE activity and inhibitors of PDEs have been used effectively as anti-inflammatory, antihypertensive, and antithrombotic agents. (Verghese, M.W. et al. (1995) Mol. Pharmacol. 47:1164-1171; and Banner, K.H.. and Page, C.P. (1995) Eur. Respir. J. 8:996-1000.)

Phospholipases (PLs) are enzymes that catalyze the removal of fatty acid residues from phosphoglycerides. PLs play an important role in transmembrane signal transduction and are named according to the specific ester bond in phosphoglycerides that is hydrolyzed, i.e., A<sub>1</sub>, A<sub>2</sub>, C or D. PLA<sub>2</sub> cleaves the ester bond at position 2 of the glycerol moiety of membrane phospholipids giving rise to arachidonic acid. Arachidonic acid is the common precursor to four major classes of eicosanoids; prostaglandins, prostacyclins, thromboxanes and leukotrienes. Eicosanoids are signaling molecules involved in the contraction of smooth muscle, platelet aggregation, and pain and inflammatory responses. PLC is an important link in certain receptor-mediated, signaling transduction pathways. Extracellular signaling molecules including hormones, growth factors, neurotransmitters, and immunoglobulins bind to their respective cell surface receptors and activate PLC. Activated PLC generates second messenger molecules from the hydrolysis of inositol phospholipids that regulate cellular processes, e.g., secretion, neural activity, metabolism and proliferation. (Alberts, B. et al. (1994) Molecular Biology of The Cell, Garland Publishing, Inc., New York, NY, pp. 85, 211, 239-240, 642-645.)

The nucleotide cyclases, i.e., adenylate and guanylate cyclase, catalyze the synthesis of the cyclic nucleotides, cAMP and cGMP, from ATP and GTP, respectively. They act in concert with phosphodiesterases, which degrade cAMP and cGMP, to regulate the cellular levels of these molecules and their functions. cAMP and cGMP function as intracellular second messengers to transduce a variety of extracellular signals, e.g., hormones, and light and neurotransmitters. Adenylate cyclase is a plasma membrane protein that is coupled with various hormone receptors also located on the plasma membrane. Binding of a hormone to its receptor activates adenylate cyclase which, in turn, increases the levels of cAMP in the cytosol. The activation of other molecules by cAMP leads to the cellular effect of the

25

30

10

5

### PF-0459 US

hormone. In a similar manner, guanylate cyclase participates in the process of visual excitation and phototransduction in the eye. (Stryer, L. (1988) Biochemistry W.H. Freeman and Co., New York, pp. 975-980, 1029-1035.) Cytokines are produced in response to cell perturbation. Some cytokines are produced as precursor forms, and some form multimers in order to become active. They are produced in groups and in patterns characteristic of the particular stimulus or disease, and the members of the group interact with one another and other molecules to produce an overall biological response. Interleukins, neurotrophins, growth factors, interferons, and chemokines are all families of cytokines which work in conjunction with cellular receptors to regulate cell proliferation and differentiation and to affect such activities, e.g., leukocyte migration and function, hematopoietic cell proliferation, temperature regulation, acute response to infections, tissue remodeling, and cell survival. Studies using antibodies or other drugs that modify the activity of a particular cytokine are used to elucidate the roles of individual cytokines in pathology and physiology.

Chemokines are a small chemoattractant cytokines which are active in leukocyte trafficking. Initially, chemokines were isolated and purified from inflamed tissues, but recently several chemokines have been discovered through molecular cloning techniques. Chemokines have been shown to be active in cell activation and migration, angiogenic and angiostatic activities, suppression of hematopoiesis, HIV infectivity, and promoting Th-1(IL-2-, interferon  $\gamma$ -stimulated) cytokine release.

Chemokines generally contain 70-100 amino acids and are subdivided into four subfamilies based on the presence and arrangement of conserved CXC, CC, CX3C and C motifs. The CXC (alpha), CC (beta), and CX3C chemokines contain four conserved cysteines. The CC subfamily is active on monocytes, lymphocytes, eosinophils, and mast cells; the CXC subfamily, on neutrophils; CX3C and C subfamilies, on T-cells. Many of the CC chemokines have been characterized functionally as well as structurally. (Callard, R. and Gearing, A. (1994) The Cytokine Facts Book, Academic Press, New York, NY, pp. 181-190, 210-213, 223-227.)

Growth and differentiation factors function in intercellular communication. Once secreted from the cell, some factors require oligomerization or association with ECM in order to function. Complex interactions among these factors and their receptors result in the

30

5

10

### PF-0459 US

stimulation or inhibition of cell division, cell differentiation, cell signaling, and cell motility. Some factors act on their cell of origin (autocrine signaling); on neighboring cells (paracrine signaling); or on distant cells (endocrine signaling).

There are three broad classes of growth and differentiation factors. The first class includes the large polypeptide growth factors, e.g., epidermal growth factor, fibroblast growth factor, transforming growth factor, insulin-like growth factor, and platelet-derived growth factor. Each of these defines a family of related molecules which stimulate cell proliferation for wound healing, bone synthesis and remodeling, and regeneration of epithelial, epidermal, and connective tissues, and induce differentiation of embryonic tissues. Nerve growth factor functions specifically as a neurotrophic factor, and all induce differentiation of embryonic tissues. The second class includes the hematopoietic growth factors which stimulate the proliferation and differentiation of blood cells such as B-lymphocytes, T-lymphocytes, erythrocytes, platelets, eosinophils, basophils, neutrophils, macrophages, and their stem cell precursors. These factors include colony-stimulating factors, erythropoietin, and cytokines, e.g., interleukins, interferons (IFNs), and tumor necrosis factor (TNF). Cytokines are secreted by cells of the immune system and function in immunomodulation. The third class includes small peptide factors e.g., bombesin, vasopressin, oxytocin, endothelin, transferrin, angiotensin II, vasoactive intestinal peptide, and bradykinin, which function as hormones to regulate cellular functions other than proliferation.

Growth and differentiation factors have been shown to play critical roles in neoplastic transformation of cells <u>in vitro</u> and in tumor progression <u>in vivo</u>. Inappropriate expression of growth factors by tumor cells may contribute to vascularization and metastasis of melanotic tumors. In hematopoiesis, growth factor misregulation can result in anemias, leukemias and lymphomas. Certain growth factors, e.g., IFN, are cytotoxic to tumor cells both <u>in vivo</u> and <u>in vitro</u>. Moreover, growth factors and/or their receptors are related both structurally and functionally related to oncoproteins. In addition, growth factors affect transcriptional regulation of both proto-oncogenes and oncosuppressor genes. (Pimentel, E. (1994) Handbook of Growth Factors, CRC Press, Ann Arbor, MI, pp. 6-25.)

Proteolytic enzymes or proteases degrade proteins by reducing the activation energy needed for the hydrolysis of peptide bonds. The major families are the zinc, serine, cysteine,

25

30

10

5

PF-0459 US

thiol, and carboxyl proteases.

Zinc proteases, e.g., carboxypeptidase A, have a zinc ion bound to the active site. recognize C-terminal residues that contain an aromatic or bulky aliphatic side chain, and hydrolyze the peptide bond adjacent to the C-terminal residues. Serine proteases have an active site serine residue and include digestive enzymes, e.g., trypsin and chymotrypsin, components of the complement and blood-clotting cascades, and enzymes that control the degradation and turnover of extracellular matrix (ECM) molecules. Subfamilies of serine proteases include tryptases (cleavage after arginine or lysine), aspases (cleavage after aspartate), chymases (cleavage after phenylalanine or leucine), metases (cleavage after methionine), and serases (cleavage after serine). Cysteine proteases (e.g. cathepsin) are produced by monocytes, macrophages and other immune cells and are involved in diverse cellular processes ranging from the processing of precursor proteins to intracellular degradation. Overproduction of these enzymes can cause the tissue destruction associated with rheumatoid arthritis and asthma. Thiol proteases, e.g., papain, contain an active site cysteine and are widely distributed within tissues. Thiol proteases effect catalysis through a thiol ester intermediate facilitated by a proximal histidine side chain. Carboxyl proteases, e.g., pepsin, are active only under acidic conditions (pH 2 to 3). The active site of pepsin contains two aspartate residues; when one aspartate is ionized and the other is not, the enzyme is active. A common feature of the carboxyl proteases is that they are inhibited by very low concentrations (10<sup>-10</sup> M) of the inhibitor pepstatin. A substrate analog which induces structural changes at the active site of a protease functions as an antagonist or inhibitor.

Guanosine triphosphate-binding proteins (G proteins) participate in intracellular signal transduction and control regulatory pathways through cell surface receptors. These receptors respond to hormones, growth factors, neuromodulators, or other signaling molecules, by binding GTP. Binding of GTP leads to the production of cAMP which controls phosphorylation and activation of other proteins. During this process, the hydrolysis of GTP acts as an energy source as well as an on-off switch for the GTPase activity.

The G proteins are small proteins which consist of single 21-30 kDa polypeptides. They can be classified into five subfamilies: Ras, Rho, Ran, Rab, and ADP-ribosylation

30

5

10

### PF-0459 US

factor. These proteins regulate cell growth, cell cycle control, protein secretion, and intracellular vesicle interaction. In particular, the Ras proteins are essential in transducing signals from receptor tyrosine kinases to serine/threonine kinases which control cell growth and differentiation. Mutant Ras proteins, which bind but can not hydrolyze GTP, are permanently activated and cause continuous cell proliferation or cancer.

All five subfamilies share common structural features and four conserved motifs, I to IV. Motif I is the most variable and has the signature of GXXXXGK, in which lysine interacts with the  $\beta$ - and  $\gamma$ -phosphate groups of GTP. Motif II, III, and IV have DTAGQE, NKXD, and EXSAX as their respective signatures and regulate the binding of g-phosphate, GTP, and the guanine base of GTP, respectively. Most of the membrane-bound G proteins require a carboxy terminal isoprenyl group (CAAX), added posttranslationally, for membrane association and biological activity. The G proteins also have a variable effector region, located between motifs I and II, which is characterized as the interaction site for guanine nucleotide exchange factors or GTPase-activating proteins.

Eukaryotic cells are bound by a membrane and subdivided into membrane bound compartments. As membranes are impermeable to many ions and polar molecules, transport of these molecules is mediated by ion channels, ion pumps, transport proteins, or pumps. Symporters and antiporters regulate cytosolic pH by transporting ions and small molecules, e.g., amino acids, glucose, and drugs, across membranes; symporters transport small molecules and ions in the same direction, and antiporters, in the opposite direction.

Transporter superfamilies include facilitative transporters and active ATP binding cassette transporters involved in multiple-drug resistance and the targeting of antigenic peptides to MHC Class I molecules. These transporters bind to a specific ion or other molecule and undergo conformational changes in order to transfer the ion or molecule across a membrane. Transport can occur by a passive, concentration-dependent mechanism or can be linked to an energy source such as ATP hydrolysis or an ion gradient.

Ion channels are formed by transmembrane proteins which form a lined passageway across the membrane through which water and ions, e.g., Na<sup>+</sup>, K<sup>+</sup>, Ca<sup>2+</sup>, and Cl<sup>-</sup>, enter and exit the cell. For example, chloride channels are involved in the regulation of the membrane electric potential as well as absorption and secretion of ions across the membrane. In

### PF-0459 US

5

10

15 the second se

intracellular membranes of the Golgi apparatus and endocytic vesicles, chloride channels also regulate organelle pH. Electrophysiological and pharmacological studies suggest that a variety of chloride channels exist in different cell types and that many of these channels have one or more protein kinase phosphorylation sites.

Ion pumps are ATPases which actively maintain membrane gradients. Ion pumps can be grouped into three classes, e.g., P, V, and F, according to their structure and function. All have one or more binding sites for ATP on the cytosolic face of the membrane. The P-class ion pumps consist of two  $\alpha$  and two  $\beta$  transmembrane subunits, include  $Ca^{2+}$  ATPase and  $Na^+/K^+$  ATPase, and function in transporting  $H^+$ ,  $Na^+$ ,  $K^+$ , and  $Ca^{2+}$  ions. The V- and F-class ion pumps have similar structures, a cytosolic domain formed by at least five extrinsic polypeptides and at least 2 transmembrane proteins, and only transport  $H^+$ . F class  $H^+$  pumps have been identified from the membranes of mitochondria and chloroplast, and V-class  $H^+$  pumps regulate acidity inside lysosomes, endosomes, and plant vacuoles.

A family of structurally related intrinsic membrane proteins known as facilitative glucose transporters catalyze the movement of glucose and other selected sugars across the plasma membrane. The proteins in this family contain a highly conserved, large transmembrane domain made of 12 transmembrane  $\alpha$ -helices, and several less conserved, asymmetric, cytoplasmic and exoplasmic domains. (Pessin, J. E., and Bell, G.I. (1992) Annu. Rev. Physiol. 54:911-930.)

Amino acid transport is mediated by Na<sup>+</sup> dependent amino acid transporters. These transporters are involved in gastrointestinal and renal uptake of dietary and cellular amino acids and the re-uptake of neurotransmitters. Transport of cationic amino acids is mediated by the system y+ family members and the cationic amino acid transporter (CAT) family. Members of the CAT family share a high degree of sequence homology, and each contains 12-14 putative transmembrane domains. (Ito, K. and Groudine, M. (1997) J. Biol. Chem. 272:26780-26786.)

Proton-coupled, 12 membrane-spanning domain transporters such as PEPT 1 and PEPT 2 are responsible for gastrointestinal absorption and for renal reabsorbtion of peptides using an electrochemical H<sup>+</sup> gradient as the driving force. A heterodimeric peptide transporter, consisting of TAP 1 and TAP 2, is associated with antigen processing. Peptide

25

the control of the co

25

30

5

10

antigens are transported across the membrane of the endoplasmic reticulum so they can be presented to the major histocompatibility complex class I molecules. Each TAP protein consists of multiple hydrophobic membrane spanning segments and a highly conserved ATP-binding cassette. (Boll, M. et al. (1996) Proc. Natl. Acad. Sci. 93:284-289.)

Hormones are secreted molecules that circulate in the body fluids and bind to specific receptors on the surface of, or within, target tissue cells. Although they have diverse biochemical compositions and mechanisms of action, hormones can be grouped into two categories. One category consists of small lipophilic molecules that diffuse through the plasma membrane of target cells, bind to cytosolic or nuclear receptors, and form a complex alters gene expression. Examples of this category include retinoic acid, thyroxine, and the cholesterol derived steroid hormones, progesterone, estrogen, testosterone, cortisol, and aldosterone. These hormones have a long half-life, e.g., several hours to days, and long-term effects of their target cells. Their solubility in the blood may be increased by their association with carrier molecules. Within the target cell nucleus, hormone/receptor complexes bind to specific response elements in target gene regulatory regions.

A second category consists of hydrophilic hormones that function by binding to cell surface receptors and transducing the signal across the plasma membrane. Examples of this category include amino acid derivatives, such as catecholamines, e.g., epinephrine, norepinephrine, and histamine; peptide hormones, e.g., glucagon, insulin, gastrin, secretin, cholecystokinin, adrenocorticotropic hormone, follicle stimulating hormone, luteinizing hormone, thyroid stimulating hormone, parathormone, and vasopressin. Peptide hormones are synthesized as inactive forms and stored in secretory vesicles. These hormones are activated by protease cleavage before being released from the cell. Many hydrophilic hormones have a very short half-life and effect, e.g., seconds to hours, and are inactivated by proteases in the blood. (Lodish et al. (1995) Molecular Cell Biology, Scientific American Books Inc., New York, NY, pp. 856-864.)

Neuropeptides and vasomediators (NP/VM) comprise a large family of endogenous signaling molecules. Included in the family are neurotransmitters such as bombesin, neuropeptide Y, neurotensin, neuromedin N, melanocortins, opioids, e.g., enkephalins, endorphins and dynorphins, galanin, somatostatin, tachykinins, vasopressin, and vasoactive

30

5

10

intestinal peptide, and circulatory system-borne signaling molecules, e.g., angiotensin, complement, calcitonin, endothelins, formyl-methionyl peptides, glucagon, cholecystokinin and gastrin. These proteins are synthesized as "pre-pro" molecules, and are activated and inactivated by proteolytic cleavage. NP/VMs can transduce signals directly, modulate the activity or release of other neurotransmitters and hormones, and act as catalytic enzymes in cascades. The effects of NP/VMs range from extremely brief or long-lasting (melanocortin-Regulatory molecules turn individual genes or mediated changes in skin melanin). groups of genes on and off in response to various inductive mechanisms of the cell or organism; act as transcription factors by determining whether or not transcription is initiated, enhanced, or repressed; and splice transcripts as dictated in a particular cell or tissue. Although they interact with short stretches of DNA scattered throughout the entire genome, most gene expression is regulated near the site at which transcription starts or within the open reading frame of the gene being expressed. The regulated stretches of the DNA can be simple and interact with only a single protein, or they can require several proteins acting as part of a complex to regulate gene expression. The external features of the double helix which provide recognition sites are hydrogen bond donor and acceptor groups, hydrophobic patches, major and minor grooves, and regular, repeated stretches of sequences which cause distinct bends in the helix. The surface features of the regulatory molecule are complementary to those of the DNA.

Many of the transcription factors incorporate one of a set of DNA-binding structural motifs, each of which contains either α helices or β sheets and binds to the major groove of DNA. Seven of the structural motifs common to transcription factors are helix-turn-helix, homeodomains, zinc finger, steroid receptor, β sheets, leucine zipper, and helix-loop-helix. (Pabo, C.O. and R.T. Sauer (1992) Ann. Rev. Biochem. 61:1053-95.) Other domains of transcription factors may form crucial contacts with the DNA. In addition, accessory proteins provide important interactions which may convert a particular protein complex to an activator or a repressor or may prevent binding. (Alberts, B. et al. (1994) Molecular Biology of the Cell, Garland Publishing Co, New York, NY pp. 401-474.)

The discovery of new human signal peptide-containing proteins and the polynucleotides encoding these molecules satisfies a need in the art by providing new

25

30

5

10

PF-0459 US

compositions which are useful in the diagnosis, treatment, and prevention of cancer and immunological disorders.

### SUMMARY OF THE INVENTION

The invention features a substantially purified human signal peptide-containing protein (SIGP), having an amino acid sequence selected from the group consisting of SEQ ID NO:1 SEQ ID NO:2, SEQ ID NO:3, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:6, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:10, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, SEQ ID NO:14, SEQ ID NO:15, SEQ ID NO:16, SEQ ID NO:17, SEQ ID NO:18, SEQ ID NO:19, SEQ ID NO:20, SEQ ID NO:21, SEQ ID NO:22, SEQ ID NO:23, SEQ ID NO:24, SEQ ID NO:25, SEQ ID NO:26, SEQ ID NO:27, SEQ ID NO:28, SEQ ID NO:29, SEQ ID NO:30, SEQ ID NO:31, SEQ ID NO:32, SEQ ID NO:33, SEQ ID NO:34, SEQ ID NO:35, SEQ ID NO:36, SEQ ID NO:37, SEQ ID NO:38, SEQ ID NO:39, SEQ ID NO:40, SEQ ID NO:41, SEQ ID NO:42, SEQ ID NO:43, SEQ ID NO:44, SEQ ID NO:45, SEQ ID NO:46, SEQ ID NO:47, SEQ ID NO:48, SEQ ID NO:49, SEQ ID NO:50, SEQ ID NO:51, SEQ ID NO:52, SEQ ID NO:53, SEQ ID NO:54, SEQ ID NO:55, SEQ ID NO:56, SEQ ID NO:57, SEQ ID NO:58, SEQ ID NO:59, SEQ ID NO:60, SEQ ID NO:61, SEQ ID NO:62, SEQ ID NO:63, SEQ ID NO:64, SEQ ID NO:65, SEQ ID NO:66, SEQ ID NO:67, SEQ ID NO:68, SEQ ID NO:69, SEQ ID NO:70, SEQ ID NO:71, SEQ ID NO:72, SEQ ID NO:73, SEQ ID NO:74, SEQ ID NO:75, SEQ ID NO:76, and SEQ ID NO:77...

The invention further provides isolated and substantially purified polynucleotides encoding SIGP. In a particular aspect, the polynucleotide has a nucleic acid sequence selected from the group consisting of SEQ ID NO:78, SEQ ID NO:79, SEQ ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEQ ID NO:85, SEQ ID NO:86, SEQ ID NO:87, SEQ ID NO:88, SEQ ID NO:89, SEQ ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID NO:115, SEQ ID NO:116, SEQ ID NO:117, SEQ ID

### PF-0459 US

5

10

The left first that the table that the state and the state with

20

25

30

NO:118, SEQ ID NO:119, SEQ ID NO:120, SEQ ID NO:121, SEQ ID NO:122, SEQ ID NO:123, SEQ ID NO:124, SEQ ID NO:125, SEQ ID NO:126, SEQ ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:132, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139, SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154.

In addition, the invention provides a polynucleotide, or fragment thereof, which hybridizes to any of the polynucleotides encoding an SIGP selected from the group consisting of SEQ ID NO:1, SEQ ID NO:2, SEQ ID NO:3, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:6, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:10, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, SEQ ID NO:14, SEQ ID NO:15, SEQ ID NO:16, SEQ ID NO:17, SEQ ID NO:18, SEQ ID NO:19, SEQ ID NO:20, SEQ ID NO:21, SEQ ID NO:22, SEQ ID NO:23, SEQ ID NO:24, SEQ ID NO:25, SEQ ID NO:26, SEQ ID NO:27, SEQ ID NO:28, SEQ ID NO:29, SEQ ID NO:30, SEQ ID NO:31, SEQ ID NO:32, SEQ ID NO:33, SEQ ID NO:34, SEQ ID NO:35, SEQ ID NO:36, SEQ ID NO:37, SEQ ID NO:38, SEQ ID NO:39, SEO ID NO:40, SEQ ID NO:41, SEQ ID NO:42, SEQ ID NO:43, SEQ ID NO:44, SEQ ID NO:45, SEQ ID NO:46, SEQ ID NO:47, SEQ ID NO:48, SEQ ID NO:49, SEQ ID NO:50, SEQ ID NO:51, SEQ ID NO:52, SEQ ID NO:53, SEQ ID NO:54, SEQ ID NO:55, SEQ ID NO:56, SEQ ID NO:57, SEQ ID NO:58, SEQ ID NO:59, SEQ ID NO:60, SEQ ID NO:61, SEQ ID NO:62, SEQ ID NO:63, SEQ ID NO:64, SEQ ID NO:65, SEQ ID NO:66, SEQ ID NO:67, SEQ ID NO:68, SEQ ID NO:69, SEQ ID NO:70, SEQ ID NO:71, SEQ ID NO:72, SEO ID NO:73, SEQ ID NO:74, SEQ ID NO:75, SEQ ID NO:76, and SEQ ID NO:77. In another aspect, the invention provides a composition comprising isolated and purified polynucleotides selected from the group consisting of SEQ ID NO:78, SEQ ID NO:79, SEQ ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEQ ID NO:85, SEQ ID NO:86, SEQ ID NO:87, SEQ ID NO:88, SEQ ID NO:89, SEQ ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID

15

30

5

10

### PF-0459 US

NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID NO:115, SEQ ID NO:116, SEQ ID NO:117, SEQ ID NO:118, SEQ ID NO:119, SEQ ID NO:120, SEQ ID NO:121, SEQ ID NO:122, SEQ ID NO:123, SEQ ID NO:124, SEQ ID NO:125, SEQ ID NO:126, SEQ ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:132, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139, SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154, or a fragment thereof.

The invention further provides a polynucleotide comprising the complement, or fragments thereof, of any one of the polynucleotides encoding SIGP. In another aspect, the invention provides compositions comprising isolated and purified polynucleotides comprising the complement of SEQ ID NO:78, SEQ ID NO:79, SEQ ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEQ ID NO:85, SEQ ID NO:86, SEQ ID NO:87, SEQ ID NO:88, SEQ ID NO:89, SEQ ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID NO:115, SEQ ID NO:116, SEQ ID NO:117, SEQ ID NO:118, SEQ ID NO:119, SEQ ID NO:120, SEQ ID NO:121, SEQ ID NO:122, SEQ ID NO:123, SEQ ID NO:124, SEQ ID NO:125, SEQ ID NO:126, SEQ ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:132, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139, SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154, or fragments thereof.

### PF-0459 US

The present invention further provides an expression vector containing at least a fragment of any one of the polynucleotides selected from the group consisting of SEQ ID NO:78, SEQ ID NO:79, SEQ ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEQ ID NO:85, SEQ ID NO:86, SEQ ID NO:87, SEQ ID NO:88, SEQ ID NO:89, SEQ ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID  $\operatorname{NO:}115,\operatorname{SEQ}\operatorname{ID}\operatorname{NO:}116,\operatorname{SEQ}\operatorname{ID}\operatorname{NO:}117,\operatorname{SEQ}\operatorname{ID}\operatorname{NO:}118,\operatorname{SEQ}\operatorname{ID}\operatorname{NO:}119,\operatorname{SEQ}\operatorname{ID}$ NO:120, SEQ ID NO:121, SEQ ID NO:122, SEQ ID NO:123, SEQ ID NO:124, SEQ ID NO:125, SEQ ID NO:126, SEQ ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:132, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139, SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154. In yet another aspect, the expression vector containing the polynucleotide is contained within a host cell.

The invention also provides a method for producing a polypeptide or a fragment thereof, the method comprising the steps of: (a) culturing the host cell containing an expression vector containing at least a fragment of a polynucleotide encoding SIGP under conditions suitable for the expression of the polypeptide; and (b) recovering the polypeptide from the host cell culture.

The invention also provides a pharmaceutical composition comprising a substantially purified SIGP in conjunction with a suitable pharmaceutical carrier.

The invention further includes a purified antibody which binds to SIGP, as well as a purified agonist and a purified antagonist of SIGP.

The invention also provides a method for treating or preventing a cancer associated with the decreased expression or activity of SIGP, the method comprising the step of

5

The series of th

25

25

30

5

10

administering to a subject in need of such treatment an effective amount of a pharmaceutical composition containing SIGP.

The invention also provides a method for treating or preventing a cancer associated with the increased expression or activity of SIGP, the method comprising the step of administering to a subject in need of such treatment an effective amount of an antagonist of SIGP.

The invention also provides a method for treating or preventing an immune response associated with the increased expression or activity of SIGP, the method comprising the step of administering to a subject in need of such treatment an effective amount of an antagonist of SIGP.

The invention also provides a method for detecting a nucleic acid sequence which encodes a human regulatory proteins in a biological sample, the method comprising the steps of: a) hybridizing a nucleic acid sequence of the biological sample to a polynucleotide sequence complementary to the polynucleotide encoding SIGP, thereby forming a hybridization complex; and b) detecting the hybridization complex, wherein the presence of the hybridization complex correlates with the presence of the nucleic acid sequence encoding the human regulatory protein in the biological sample.

The invention also provides a microarray containing at least a fragment of at least one of the polynucleotides encoding a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:1, SEQ ID NO:2, SEQ ID NO:3, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:6, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:10, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, SEQ ID NO:14, SEQ ID NO:15, SEQ ID NO:16, SEQ ID NO:17, SEQ ID NO:18, SEQ ID NO:19, SEQ ID NO:20, SEQ ID NO:21, SEQ ID NO:22, SEQ ID NO:23, SEQ ID NO:24, SEQ ID NO:25, SEQ ID NO:26, SEQ ID NO:27, SEQ ID NO:28, SEQ ID NO:29, SEQ ID NO:30, SEQ ID NO:31, SEQ ID NO:32, SEQ ID NO:33, SEQ ID NO:34, SEQ ID NO:35, SEQ ID NO:36, SEQ ID NO:37, SEQ ID NO:38, SEQ ID NO:39, SEQ ID NO:40, SEQ ID NO:41, SEQ ID NO:42, SEQ ID NO:43, SEQ ID NO:44, SEQ ID NO:45, SEQ ID NO:45, SEQ ID NO:45, SEQ ID NO:51, SEQ ID NO:52, SEQ ID NO:53, SEQ ID NO:54, SEQ ID NO:55, SEQ ID NO:55, SEQ ID NO:55, SEQ ID NO:56, SEQ ID NO:57, SEQ ID NO:58, SEQ ID NO:59, SEQ ID NO:60,

25

30

5

PF-0459 US

SEQ ID NO:61, SEQ ID NO:62, SEQ ID NO:63, SEQ ID NO:64, SEQ ID NO:65, SEQ ID NO:66, SEQ ID NO:67, SEQ ID NO:68, SEQ ID NO:69, SEQ ID NO:70, SEQ ID NO:71, SEQ ID NO:72, SEQ ID NO:73, SEQ ID NO:74, SEQ ID NO:75, SEQ ID NO:76, and SEQ ID NO:77.

The invention also provides a method for detecting the expression level of a nucleic acid encoding a human regulatory protein in a biological sample, the method comprising the steps of hybridizing the nucleic acid sequence of the biological sample to a complementary polynucleotide, thereby forming hybridization complex; and determining expression of the nucleic acid sequence encoding a human regulatory protein in the biological sample by identifying the presence of the hybridization complex. In a preferred embodiment, prior to the hybridizing step, the nucleic acid sequences of the biological sample are amplified and labeled by the polymerase chain reaction.

### **DESCRIPTION OF THE INVENTION**

Before the present proteins, nucleotide sequences, and methods are described, it is understood that this invention is not limited to the particular methodology, protocols, cell lines, vectors, and reagents described, as these may vary. It is also to be understood that the terminology used herein is for the purpose of describing particular embodiments only, and is not intended to limit the scope of the present invention which will be limited only by the appended claims.

It must be noted that as used herein and in the appended claims, the singular forms "a," "an," and "the" include plural reference unless the context clearly dictates otherwise. Thus, for example, a reference to "a host cell" includes a plurality of such host cells, and a reference to "an antibody" is a reference to one or more antibodies and equivalents thereof known to those skilled in the art, and so forth.

Unless defined otherwise, all technical and scientific terms used herein have the same meanings as commonly understood by one of ordinary skill in the art to which this invention belongs. Although any methods and materials similar or equivalent to those described herein can be used in the practice or testing of the present invention, the preferred methods, devices, and materials are now described. All publications mentioned herein are cited for the purpose

10

If the first state that the state state is the state of t

25

30

of describing and disclosing the cell lines, vectors, and methodologies which are reported in the publications and which might be used in connection with the invention. Nothing herein is to be construed as an admission that the invention is not entitled to antedate such disclosure by virtue of prior invention.

### **DEFINITIONS**

"SIGP," as used herein, refers to the amino acid sequences of substantially purified SIGP obtained from any species, particularly a mammalian species, including bovine, ovine, porcine, murine, equine, and preferably the human species, from any source, whether natural, synthetic, semi-synthetic, or recombinant.

The term "agonist," as used herein, refers to a molecule which, when bound to SIGP, increases or prolongs the duration of the effect of SIGP. Agonists may include proteins, nucleic acids, carbohydrates, or any other molecules which bind to and modulate the effect of SIGP.

An "allele" or an "allelic sequence," as these terms are used herein, is an alternative form of the gene encoding SIGP. Alleles may result from at least one mutation in the nucleic acid sequence and may result in altered mRNAs or in polypeptides whose structure or function may or may not be altered. Any given natural or recombinant gene may have none, one, or many allelic forms. Common mutational changes which give rise to alleles are generally ascribed to natural deletions, additions, or substitutions of nucleotides. Each of these types of changes may occur alone, or in combination with the others, one or more times in a given sequence.

"Altered" nucleic acid sequences encoding SIGP, as described herein, include those sequences with deletions, insertions, or substitutions of different nucleotides, resulting in a polynucleotide the same SIGP or a polypeptide with at least one functional characteristic of SIGP. Included within this definition are polymorphisms which may or may not be readily detectable using a particular oligonucleotide probe of the polynucleotide encoding SIGP, and improper or unexpected hybridization to alleles, with a locus other than the normal chromosomal locus for the polynucleotide sequence encoding SIGP. The encoded protein may also be "altered," and may contain deletions, insertions, or substitutions of amino acid

Some the state of the state of

25

30

5

10

residues which produce a silent change and result in a functionally equivalent SIGP. Deliberate amino acid substitutions may be made on the basis of similarity in polarity, charge, solubility, hydrophobicity, hydrophilicity, and/or the amphipathic nature of the residues, as long as the biological or immunological activity of SIGP is retained. For example, negatively charged amino acids may include aspartic acid and glutamic acid, positively charged amino acids may include lysine and arginine, and amino acids with uncharged polar head groups having similar hydrophilicity values may include leucine, isoleucine, and valine; glycine and alanine; asparagine and glutamine; serine and threonine; and phenylalanine and tyrosine.

The terms "amino acid" or "amino acid sequence," as used herein, refer to an oligopeptide, peptide, polypeptide, or protein sequence, or a fragment of any of these, and to naturally occurring or synthetic molecules. In this context, "fragments", "immunogenic fragments", or "antigenic fragments" refer to fragments of SIGP which are preferably about 5 to about 15 amino acids in length and which retain some biological activity or immunological activity of SIGP. Where "amino acid sequence" is recited herein to refer to an amino acid sequence of a naturally occurring protein molecule, "amino acid sequence" and like terms are not meant to limit the amino acid sequence to the complete native amino acid sequence associated with the recited protein molecule.

"Amplification," as used herein, relates to the production of additional copies of a nucleic acid sequence. Amplification is generally carried out using polymerase chain reaction (PCR) technologies well known in the art. (See, e.g., Dieffenbach, C.W. and G.S. Dveksler (1995) PCR Primer, a Laboratory Manual, Cold Spring Harbor Press, Plainview, NY, pp.1-5.)

The term "antagonist," as it is used herein, refers to a molecule which, when bound to SIGP, decreases the amount or the duration of the effect of the biological or immunological activity of SIGP. Antagonists may include proteins, nucleic acids, carbohydrates, antibodies, or any other molecules which decrease the effect of SIGP.

As used herein, the term "antibody" refers to intact molecules as well as to fragments thereof, such as Fa, F(ab')<sub>2</sub>, and Fv fragments, which are capable of binding the epitopic determinant. Antibodies that bind SIGP polypeptides can be prepared using intact polypeptides or using fragments containing small peptides of interest as the immunizing

; <sub>10</sub>

The last that the time is a way of the time with all last the second with a last time.

antigen. The polypeptide or oligopeptide used to immunize an animal (e.g., a mouse, a rat, or a rabbit) can be derived from the translation of RNA, or synthesized chemically, and can be conjugated to a carrier protein if desired. Commonly used carriers that are chemically coupled to peptides include bovine serum albumin, thyroglobulin, and keyhole limpet hemocyanin (KLH). The coupled peptide is then used to immunize the animal.

The term "antigenic determinant," as used herein, refers to that fragment of a molecule (i.e., an epitope) that makes contact with a particular antibody. When a protein or a fragment of a protein is used to immunize a host animal, numerous regions of the protein may induce the production of antibodies which bind specifically to antigenic determinants (given regions or three-dimensional structures on the protein). An antigenic determinant may compete with the intact antigen (i.e., the immunogen used to elicit the immune response) for binding to an antibody.

The term "antisense," as used herein, refers to any composition containing a nucleic acid sequence which is complementary to a specific nucleic acid sequence. The term "antisense strand" is used in reference to a nucleic acid strand that is complementary to the "sense" strand. Antisense molecules may be produced by any method including synthesis or transcription. Once introduced into a cell, the complementary nucleotides combine with natural sequences produced by the cell to form duplexes and to block either transcription or translation. The designation "negative" can refer to the antisense strand, and the designation "positive" can refer to the sense strand.

As used herein, the term "biologically active," refers to a protein having structural, regulatory, or biochemical functions of a naturally occurring molecule. Likewise, "immunologically active" refers to the capability of the natural, recombinant, or synthetic SIGP, or of any oligopeptide thereof, to induce a specific immune response in appropriate animals or cells and to bind with specific antibodies.

The terms "complementary" or "complementarity," as used herein, refer to the natural binding of polynucleotides under permissive salt and temperature conditions by base pairing. For example, the sequence "A-G-T" binds to the complementary sequence "T-C-A." Complementarity between two single-stranded molecules may be "partial," such that only some of the nucleic acids bind, or it may be "complete," such that total complementarity

25

30

The first first first such that the set of the such that we will refer the first such that the set of the such that the set of the such that we will refer the such that w

25

30

5

10

exists between the single stranded molecules. The degree of complementarity between nucleic acid strands has significant effects on the efficiency and strength of the hybridization between the nucleic acid strands. This is of particular importance in amplification reactions, which depend upon binding between nucleic acids strands, and in the design and use of peptide nucleic acid (PNA) molecules.

A "composition comprising a given polynucleotide sequence" or a "composition comprising a given amino acid sequence," as these terms are used herein, refer broadly to any composition containing the given polynucleotide or amino acid sequence. The composition may comprise a dry formulation, an aqueous solution, or a sterile composition. Compositions comprising polynucleotides encoding SIGP, e.g., SEQ ID NO:78, SEQ ID NO:79, SEQ ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEQ ID NO:85, SEQ ID NO:86, SEQ ID NO:87, SEQ ID NO:88, SEQ ID NO:89, SEQ ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID NO:115, SEQ ID NO:116, SEQ ID NO:117, SEQ ID NO:118, SEQ ID NO:119, SEQ ID NO:120, SEQ ID NO:121, SEQ ID NO:122, SEQ ID NO:123, SEQ ID NO:124, SEQ ID NO:125, SEQ ID NO:126, SEQ ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:132, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139, SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154, or fragments thereof, may be employed as hybridization probes. The probes may be stored in freeze-dried form and may be associated with a stabilizing agent such as a carbohydrate. In hybridizations, the probe may be deployed in an aqueous solution containing salts (e.g., NaCl), detergents (e.g., SDS) and other components (e.g., Denhardt's solution, dry milk, salmon sperm DNA, etc.).

The phrase "consensus sequence," as used herein, refers to a nucleic acid sequence

### PF-0459 US

which has been resequenced to resolve uncalled bases, extended using XL-PCR<sup>TM</sup> (Perkin Elmer, Norwalk, CT) in the 5' and/or the 3' direction, and resequenced, or which has been assembled from the overlapping sequences of more than one Incyte Clone using a computer program for fragment assembly, such as the GELVIEW<sup>TM</sup> Fragment Assembly system (GCG, Madison, WI). Some sequences have been both extended and assembled to produce the consensus sequence.

As used herein, the term "correlates with expression of a polynucleotide" indicates that the detection of the presence of nucleic acids, the same or related to a nucleic acid sequence encoding SIGP, by northern analysis is indicative of the presence of nucleic acids encoding SIGP in a sample, and thereby correlates with expression of the transcript from the polynucleotide encoding SIGP.

The term "SIGP" refers to any or all of the human polypeptides, SIGP-1, SIGP-2, SIGP-3, SIGP-4, SIGP-5, SIGP-6, SIGP-7, SIGP-8, SIGP-9, SIGP-10, SIGP-11, SIGP-12, SIGP-13, SIGP-14, SIGP-15, SIGP-16, SIGP-17, SIGP-18, SIGP-19, SIGP-20, SIGP-21, SIGP-22, SIGP-23, SIGP-24, SIGP-25, SIGP-26, SIGP-27, SIGP-28, SIGP-29, SIGP-30, SIGP-31, SIGP-32, SIGP-33, SIGP-34, SIGP-35, SIGP-36, SIGP-37, SIGP-38, SIGP-39, SIGP-40, SIGP-41, SIGP-42, SIGP-43, SIGP-44, SIGP-45, SIGP-46, SIGP-47, SIGP-48, SIGP-49, SIGP-50, SIGP-51, SIGP-52, SIGP-53, SIGP-54, SIGP-55, SIGP-56, SIGP-57, SIGP-58, SIGP-59, SIGP-60, SIGP-61, SIGP-62, SIGP-63, SIGP-64, SIGP-65, SIGP-66, SIGP-67, SIGP-68, SIGP-69, SIGP-70, SIGP-71, SIGP-72, SIGP-73, SIGP-74, SIGP-75, SIGP-76, and SIGP-77.

A "deletion," as the term is used herein, refers to a change in the amino acid or nucleotide sequence that results in the absence of one or more amino acid residues or nucleotides.

The term "derivative," as used herein, refers to the chemical modification of SIGP, of a polynucleotide sequence encoding SIGP, or of a polynucleotide sequence complementary to a polynucleotide sequence encoding SIGP. Chemical modifications of a polynucleotide sequence can include, for example, replacement of hydrogen by an alkyl, acyl, or amino group. A derivative polynucleotide encodes a polypeptide which retains at least one biological or immunological function of the natural molecule. A derivative polypeptide is

20

25

30

5

10

25

30

5

10

### PF-0459 US

one modified by glycosylation, pegylation, or any similar process that retains at least one biological or immunological function of the polypeptide from which it was derived.

The term "homology," as used herein, refers to a degree of complementarity. There may be partial homology or complete homology. The word "identity" may substitute for the word "homology." A partially complementary sequence that at least partially inhibits an identical sequence from hybridizing to a target nucleic acid is referred to as "substantially homologous." The inhibition of hybridization of the completely complementary sequence to the target sequence may be examined using a hybridization assay (Southern or northern blot, solution hybridization, and the like) under conditions of reduced stringency. A substantially homologous sequence or hybridization probe will compete for and inhibit the binding of a completely homologous sequence to the target sequence under conditions of reduced stringency. This is not to say that conditions of reduced stringency are such that non-specific binding is permitted, as reduced stringency conditions require that the binding of two sequences to one another be a specific (i.e., a selective) interaction. The absence of nonspecific binding may be tested by the use of a second target sequence which lacks even a partial degree of complementarity (e.g., less than about 30% homology or identity). In the absence of non-specific binding, the substantially homologous sequence or probe will not hybridize to the second non-complementary target sequence.

The phrases "percent identity" or "% identity" refer to the percentage of sequence similarity found in a comparison of two or more amino acid or nucleic acid sequences. Percent identity can be determined electronically, e.g., by using the MegAlign program (Lasergene software package, DNASTAR, Inc., Madison WI). The MegAlign program can create alignments between two or more sequences according to different methods, e.g., the Clustal Method. (Higgins, D.G. and Sharp, P.M. (1988) Gene 73:237-244.) The Clustal algorithm groups sequences into clusters by examining the distances between all pairs. The clusters are aligned pairwise and then in groups. The percentage similarity between two amino acid sequences, e.g., sequence A and sequence B, is calculated by dividing the length of sequence A, minus the number of gap residues in sequence A, minus the number of gap residues in sequence B, into the sum of the residue matches between sequence A and sequence B, times one hundred. Gaps of low or of no homology between the two amino acid

30

5

sequences are not included in determining percentage similarity. Percent identity between nucleic acid sequences can also be calculated by the Clustal Method, or by other methods known in the art, such as the Jotun Hein Method. (See, e.g., Hein, J. (1990) Methods in Enzymology 183:626-645.) Identity between sequences can also be determined by other methods known in the art, e.g., by varying hybridization conditions.

"Human artificial chromosomes" (HACs), as described herein, are linear microchromosomes which may contain DNA sequences of about 6 kb to 10 Mb in size, and which contain all of the elements required for stable mitotic chromosome segregation and maintenance. (See, e.g., Harrington, J.J. et al. (1997) Nat Genet. 15:345-355.)

The term "humanized antibody," as used herein, refers to antibody molecules in which the amino acid sequence in the non-antigen binding regions has been altered so that the antibody more closely resembles a human antibody, and still retains its original binding ability.

"Hybridization," as the term is used herein, refers to any process by which a strand of nucleic acid binds with a complementary strand through base pairing.

As used herein, the term "hybridization complex" as used herein, refers to a complex formed between two nucleic acid sequences by virtue of the formation of hydrogen bonds between complementary bases. A hybridization complex may be formed in solution (e.g.,  $C_0$ t or  $R_0$ t analysis) or formed between one nucleic acid sequence present in solution and another nucleic acid sequence immobilized on a solid support (e.g., paper, membranes, filters, chips, pins or glass slides, or any other appropriate substrate to which cells or their nucleic acids have been fixed).

The words "insertion" or "addition," as used herein, refer to changes in an amino acid or nucleotide sequence resulting in the addition of one or more amino acid residues or nucleotides, respectively, to the sequence found in the naturally occurring molecule.

"Immune response" can refer to conditions associated with inflammation, trauma, immune disorders, or infectious or genetic disease, etc. These conditions can be characterized by expression of various factors, e.g., cytokines, chemokines, and other signaling molecules, which may affect cellular and systemic defense systems.

The term "microarray," as used herein, refers to an array of distinct polynucleotides or

25

30

### PF-0459 US

oligonucleotides arrayed on a substrate, such as paper, nylon or any other type of membrane, filter, chip, glass slide, or any other suitable solid support.

The term "modulate," as it appears herein, refers to a change in the activity of SIGP. For example, modulation may cause an increase or a decrease in protein activity, binding characteristics, or any other biological, functional, or immunological properties of SIGP.

The phrases "nucleic acid" or "nucleic acid sequence," as used herein, refer to an oligonucleotide, nucleotide, polynucleotide, or any fragment thereof, to DNA or RNA of genomic or synthetic origin which may be single-stranded or double-stranded and may represent the sense or the antisense strand, to peptide nucleic acid (PNA), or to any DNA-like or RNA-like material. In this context, "fragments" refers to those nucleic acid sequences which are greater than about 60 nucleotides in length, and most preferably are at least about 100 nucleotides, at least about 1000 nucleotides, or at least about 10,000 nucleotides in length.

The terms "operably associated" or "operably linked," as used herein, refer to functionally related nucleic acid sequences. A promoter is operably associated or operably linked with a coding sequence if the promoter controls the transcription of the encoded polypeptide. While operably associated or operably linked nucleic acid sequences can be contiguous and in reading frame, certain genetic elements, e.g., repressor genes, are not contiguously linked to the encoded polypeptide but still bind to operator sequences that control expression of the polypeptide.

The term "oligonucleotide," as used herein, refers to a nucleic acid sequence of at least about 6 nucleotides to 60 nucleotides, preferably about 15 to 30 nucleotides, and most preferably about 20 to 25 nucleotides, which can be used in PCR amplification or in a hybridization assay or microarray. As used herein, the term "oligonucleotide" is substantially equivalent to the terms "amplimers," "primers," "oligomers," and "probes," as these terms are commonly defined in the art.

"Peptide nucleic acid" (PNA), as used herein, refers to an antisense molecule or anti-gene agent which comprises an oligonucleotide of at least about 5 nucleotides in length linked to a peptide backbone of amino acid residues ending in lysine. The terminal lysine confers solubility to the composition. PNAs preferentially bind complementary single

30

5

stranded DNA and RNA and stop transcript elongation, and may be pegylated to extend their lifespan in the cell. (See, e.g., Nielsen, P.E. et al. (1993) Anticancer Drug Des. 8:53-63.)

The term "sample," as used herein, is used in its broadest sense. A biological sample suspected of containing nucleic acids encoding SIGP, or fragments thereof, or SIGP itself may comprise a bodily fluid; an extract from a cell, chromosome, organelle, or membrane isolated from a cell; a cell; genomic DNA, RNA, or cDNA, in solution or bound to a solid support; a tissue; a tissue print; etc.

As used herein, the terms "specific binding" or "specifically binding" refer to that interaction between a protein or peptide and an agonist, an antibody, or an antagonist. The interaction is dependent upon the presence of a particular structure of the protein recognized by the binding molecule (i.e., the antigenic determinant or epitope). For example, if an antibody is specific for epitope "A," the presence of a polypeptide containing the epitope A, or the presence of free unlabeled A, in a reaction containing free labeled A and the antibody will reduce the amount of labeled A that binds to the antibody.

As used herein, the term "stringent conditions" refers to conditions which permit hybridization between polynucleotide sequences and the claimed polynucleotide sequences. Suitably stringent conditions can be defined by, for example, the concentrations of salt or formamide in the prehybridization and hybridization solutions, or by the hybridization temperature, and are well known in the art. In particular, stringency can be increased by reducing the concentration of salt, increasing the concentration of formamide, or raising the hybridization temperature.

For example, hybridization under high stringency conditions could occur in about 50% formamide at about 37°C to 42°C. Hybridization could occur under reduced stringency conditions in about 35% to 25% formamide at about 30°C to 35°C. In particular, hybridization could occur under high stringency conditions at 42°C in 50% formamide, 5X SSPE, 0.3% SDS, and 200  $\mu$ g/ml sheared and denatured salmon sperm DNA. Hybridization could occur under reduced stringency conditions as described above, but in 35% formamide at a reduced temperature of 35°C. The temperature range corresponding to a particular level of stringency can be further narrowed by calculating the purine to pyrimidine ratio of the nucleic acid of interest and adjusting the temperature accordingly. Variations on the above

25

30

5

ranges and conditions are well known in the art.

The term "substantially purified," as used herein, refers to nucleic acid or amino acid sequences that are removed from their natural environment and are isolated or separated, and are at least about 60% free, preferably about 75% free, and most preferably about 90% free from other components with which they are naturally associated.

A "substitution," as used herein, refers to the replacement of one or more amino acids or nucleotides by different amino acids or nucleotides, respectively.

"Transformation," as defined herein, describes a process by which exogenous DNA enters and changes a recipient cell. Transformation may occur under natural or artificial conditions according to various methods well known in the art, and may rely on any known method for the insertion of foreign nucleic acid sequences into a prokaryotic or eukaryotic host cell. The method for transformation is selected based on the type of host cell being transformed and may include, but is not limited to, viral infection, electroporation, heat shock, lipofection, and particle bombardment. The term "transformed" cells includes stably transformed cells in which the inserted DNA is capable of replication either as an autonomously replicating plasmid or as part of the host chromosome, and refers to cells which transiently express the inserted DNA or RNA for limited periods of time.

A "variant" of SIGP, as used herein, refers to an amino acid sequence that is altered by one or more amino acids. The variant may have "conservative" changes, wherein a substituted amino acid has similar structural or chemical properties (e.g., replacement of leucine with isoleucine). More rarely, a variant may have "nonconservative" changes (e.g., replacement of glycine with tryptophan). Analogous minor variations may also include amino acid deletions or insertions, or both. Guidance in determining which amino acid residues may be substituted, inserted, or deleted without abolishing biological or immunological activity may be found using computer programs well known in the art, for example, DNASTAR software.

### THE INVENTION

The invention is based on the discovery of new human signal peptide-containing proteins, collectively referred to as SIGP and individually as SIGP-1, SIGP-2, SIGP-3,

30

5

10

### PF-0459 US

SIGP-4, SIGP-5, SIGP-6, SIGP-7, SIGP-8, SIGP-9, SIGP-10, SIGP-11, SIGP-12, SIGP-13, SIGP-14, SIGP-15, SIGP-16, SIGP-17, SIGP-18, SIGP-19, SIGP-20, SIGP-21, SIGP-22, SIGP-23, SIGP-24, SIGP-25, SIGP-26, SIGP-27, SIGP-28, SIGP-29, SIGP-30, SIGP-31, SIGP-32, SIGP-33, SIGP-34, SIGP-35, SIGP-36, SIGP-37, SIGP-38, SIGP-39, SIGP-40, SIGP-41, SIGP-42, SIGP-43, SIGP-44, SIGP-45, SIGP-46, SIGP-47, SIGP-48, SIGP-49, SIGP-50, SIGP-51, SIGP-52, SIGP-53, SIGP-54, SIGP-55, SIGP-56, SIGP-57, SIGP-58, SIGP-59, SIGP-60, SIGP-61, SIGP-62, SIGP-63, SIGP-64, SIGP-65, SIGP-66, SIGP-67, SIGP-68, SIGP-69, SIGP-70, SIGP-71, SIGP-72, SIGP-73, SIGP-74, SIGP-75, SIGP-76, and SIGP-77; the polynucleotides encoding SIGP (SEQ ID NO:78, SEQ ID NO:79, SEQ ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEQ ID NO:85, SEQ ID NO:86, SEQ ID NO:87, SEQ ID NO:88, SEQ ID NO:89, SEQ ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID NO:115, SEQ ID NO:116, SEQ ID NO:117, SEQ ID NO:118, SEQ ID NO:119, SEQ ID NO:120, SEQ ID NO:121, SEQ ID NO:122, SEO ID NO:123, SEO ID NO:124, SEO ID NO:125, SEO ID NO:126, SEO ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:132, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139, SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154); and the use of these compositions for the diagnosis, treatment, or prevention of cancer and immunological disorders. Table 1 shows the sequence identification numbers, Incyte Clone identification number, cDNA library, NCBI sequence identifier and GenBank species description for each of the human signal peptide-containing proteins disclosed herein.

Nucleic acids encoding the SIGP-1 of the present invention were first identified in

## TABLE 1

NO.1         SEQ ID NO:78         305841         HEARNOT01         GI 505652           NO.2         SEQ ID NO:79         322866         EOSHHETO2         GI 180141           NO.3         SEQ ID NO:80         546656         BEPINOT01         GI 2290530           NO:4         SEQ ID NO:81         693453         SYNORAT03         GI 1488683           NO:5         SEQ ID NO:82         866885         BRAITUT03         GI 1488683           NO:6         SEQ ID NO:84         1255027         LUNGNOT03         GI 1684845           NO:7         SEQ ID NO:84         1255027         LUNGNOT03         GI 1684845           NO:8         SEQ ID NO:84         1255027         LUNGNOT02         GI 1684845           NO:9         SEQ ID NO:84         1255027         LUNGNOT12         GI 56805           NO:10         SEQ ID NO:85         1275261         TESTTUT02         GI 56805           NO:10         SEQ ID NO:89         1360501         LUNGNOT12         GI 205250           NO:11         SEQ ID NO:90         1362406         LUNGNOT12         GI 205250           NO:12         SEQ ID NO:90         1416553         BRAINOT12         GI 205250           NO:13         SEQ ID NO:90         1416553	0.040	Ninclootide	Clone ID	Library	NCBI I.D.	Homolog species
SEQ ID NO:79         322866         EOSIHETO2         GI 180141           SEQ ID NO:80         546656         BEPINOT01         GI 2290530           SEQ ID NO:81         693453         SYNORAT03         GI 1488683           SEQ ID NO:82         866885         BRAITUT03         GI 1488683           SEQ ID NO:83         1242271         LUNGNOT03         GI 1523073           SEQ ID NO:84         1255027         LUNGFETU3         GI 1684845           SEQ ID NO:85         1273453         TESTTUT02         GI 1684845           SEQ ID NO:86         1275261         TESTTUT02         GI 56805           SEQ ID NO:87         1281682         COLNNOT16         GI 1019433           SEQ ID NO:89         1360501         LUNGNOT12         GI 1019433           IS SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           IS SEQ ID NO:90         1415523         BRAINOT12         GI 205250           IS SEQ ID NO:90         1416553         BRAINOT12         GI 2145052           IS SEQ ID NO:90         1416553         BRAINOT03         GI 1515161           IS SEQ ID NO:90         1416553         BRAINOT04         GI 2145052           IS SEQ ID NO:90         1534876         SPLNNOT04		SEO ID NO.78	305841	HEARNOT01	GI 505652	Homo sapiens
SEQ ID NO:80         546656 BEPINOT01         GI 2290530           SEQ ID NO:81         693453 SYNORAT03         GI 1419461           SEQ ID NO:82         866885 BRAITUT03         GI 1488683           SEQ ID NO:83         1242271 LUNGNOT03         GI 1523073           SEQ ID NO:84         1255027 LUNGFET03         GI 1684845           SEQ ID NO:85         1273453 TESTTUT02         GI 56805           SEQ ID NO:86         1275261 TESTTUT02         GI 56805           SEQ ID NO:87         1281682 COLNNOT16         GI 56805           SEQ ID NO:89         1360501 LUNGNOT12         GI 1019433           SEQ ID NO:90         1362406 LUNGNOT12         GI 2072705           SEQ ID NO:91         1405329 LATRTUT02         GI 205250           SEQ ID NO:92         1415223 BRAINOT12         GI 205250           IS SEQ ID NO:94         1418517 KIDNNOT09         GI 1515161           IS SEQ ID NO:95         1440381 THYRNOT03         GI 165459           SEQ ID NO:96         1440381 THYRNOT04         GI 205250           SEQ ID NO:97         1510839 LUNGNOT04         GI 206567           SEQ ID NO:99         1559131 SPLNNOT04         GI 206924           SEQ ID NO:100         1601473 BLADNOT03         GI 2196924           SEQ ID NO:102 </td <td></td> <td>SEO ID NO.79</td> <td>322866</td> <td>EOSIHET02</td> <td>GI 180141</td> <td>Homo sapiens</td>		SEO ID NO.79	322866	EOSIHET02	GI 180141	Homo sapiens
SEQ ID NO:81         693453         SYNORAT03         GI 1419461           SEQ ID NO:82         866885         BRAITUT03         GI 1488683           SEQ ID NO:83         1242271         LUNGNOT03         GI 1523073           SEQ ID NO:84         1255027         LUNGRET03         GI 1684845           SEQ ID NO:85         1273261         TESTTUT02         GI 56805           SEQ ID NO:86         1275261         TESTTUT02         GI 56805           SEQ ID NO:87         1281682         COLNNOT16         GI 1019433           SEQ ID NO:89         1298305         BRSTNOT07         GI 1019433           SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           SEQ ID NO:90         1415223         BRAINOT12         GI 205250           SEQ ID NO:90         1415523         BRAINOT12         GI 1065459           SEQ ID NO:90         1416553         BRAINOT09         GI 1515161           SEQ ID NO:90         1416553         BRAINOT09         GI 1665459           SEQ ID NO:90         1416551         RPINNOT09         GI 1665459           SEQ ID NO:90         1510839         LUNGNOT09         GI 1665459           SEQ ID NO:90         1559131         SPLNNOT09         GI 2196924 <td></td> <td>SEO ID NO:80</td> <td>546656</td> <td>BEPINOT01</td> <td>GI 2290530</td> <td>Homo sapiens</td>		SEO ID NO:80	546656	BEPINOT01	GI 2290530	Homo sapiens
SEQ ID NO:82         866885         BRAITUT03         GI 1488683           SEQ ID NO:83         1242271         LUNGNOT03         GI 1523073           SEQ ID NO:84         1255027         LUNGRET03         GI 1684845           SEQ ID NO:85         1273453         TESTTUT02         GI 56805           SEQ ID NO:86         1275261         TESTTUT02         GI 56805           SEQ ID NO:87         1281682         COLNNOT16         GI 56805           SEQ ID NO:89         1360501         LUNGNOT12         GI 1019433           SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           SEQ ID NO:91         1405329         LATRTUT02         GI 205250           IS SEQ ID NO:92         1415223         BRAINOT12         GI 205250           IS SEQ ID NO:93         1416553         BRAINOT14         GI 205459           IS SEQ ID NO:94         1418517         KIDNNOT09         GI 1065459           IS SEQ ID NO:95         1438165         PANCNOT08         GI 145665           IS SEQ ID NO:99         1510839         LUNGNOT04         GI 1496667           IS SEQ ID NO:99         1559131         SPLNNOT04         GI 496667           IS SEQ ID NO:100         1601473         BLADNOT03		SEO ID NO:81	693453	SYNORAT03	GI 1419461	Caenorhabditis elegans
SEQ ID NO:83         1242271         LUNGNOT03         GI 1523073           SEQ ID NO:84         1255027         LUNGRET03         GI 1684845           SEQ ID NO:85         1273453         TESTTUT02         GI 56805           SEQ ID NO:86         1275261         TESTTUT02         GI 56805           1         SEQ ID NO:87         1281682         COLNNOT16           2         SEQ ID NO:88         1298305         BRSTNOT07           3         SEQ ID NO:90         1362406         LUNGNOT12         GI 1019433           4         SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           5         SEQ ID NO:91         1405329         LATRTUT02         GI 205250           6         SEQ ID NO:92         1416553         BRAINOT12         GI 205250           7         SEQ ID NO:94         1418517         KIDNNOT09         GI 1065459           8         SEQ ID NO:95         1440381         THYRNOT03         GI 1065459           9         SEQ ID NO:96         1440381         THYRNOT04         GI 296667           20         SEQ ID NO:99         1534876         SPLNNOT04         GI 296667           22         SEQ ID NO:99         1559131         SPLNNOT03		SEQ ID NO:82	866885	BRAITUT03	GI 1488683	Rattus norvegicus
SEQ ID NO:84         1255027         LUNGFET03         GI 1684845           SEQ ID NO:85         1273453         TESTTUT02         GI 56805           SEQ ID NO:86         1275261         TESTTUT02         GI 56805           1 SEQ ID NO:87         1281682         COLNNOT16         GI 56805           2 SEQ ID NO:88         1298305         BRSTNOT07         GI 1019433           3 SEQ ID NO:89         1360501         LUNGNOT12         GI 2072705           4 SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           5 SEQ ID NO:91         1405329         LATRTUT02         GI 205250           6 SEQ ID NO:92         1416553         BRAINOT12         GI 205250           7 SEQ ID NO:93         1416553         BRAINOT12         GI 2055250           8 SEQ ID NO:94         1418517         KIDNNOT09         GI 1065459           9 SEQ ID NO:95         1440381         THYRNOT04         GI 2145052           10 SEQ ID NO:96         1440381         THYRNOT04         GI 2065459           11 SEQ ID NO:99         1559131         SPLNNOT04         GI 206667           12 SEQ ID NO:99         1559131         SPLNNOT09         GI 2196924           12 SEQ ID NO:100         1601473         BLADNOT03 </td <td>SEO ID NO:6</td> <td>SEQ ID NO:83</td> <td>1242271</td> <td>LUNGNOT03</td> <td>GI 1523073</td> <td>Homo sapiens</td>	SEO ID NO:6	SEQ ID NO:83	1242271	LUNGNOT03	GI 1523073	Homo sapiens
SEQ ID NO:85         1273453         TESTTUT02         GI 56805           0         SEQ ID NO:87         1281682         COLNNOT16         GI 56805           1         SEQ ID NO:88         1298305         BRSTNOT07         GI 1019433           2         SEQ ID NO:89         1360501         LUNGNOT12         GI 2072705           3         SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           4         SEQ ID NO:90         1405329         LATRTUT02         GI 205250           5         SEQ ID NO:92         1415223         BRAINOT12         GI 205250           6         SEQ ID NO:93         1416553         BRAINOT12         GI 205250           7         SEQ ID NO:94         1418517         KIDNNOT09         GI 1065459           18         SEQ ID NO:95         1440381         THYRNOT03         GI 1515161           20         SEQ ID NO:96         1440381         THYRNOT04         GI 245065           21         SEQ ID NO:99         1534876         SPLNNOT04         GI 296667           22         SEQ ID NO:99         1559131         SPLNNOT03         GI 296667           23         SEQ ID NO:100         1601473         BLADNOT03         GI 2196924     <	SEO ID NO:7	SEQ ID NO:84	1255027	LUNGFET03	GI 1684845	Canis familiaris
SEQ ID NO:86         1275261         TESTTUT02         GI 56805           0         SEQ ID NO:87         1281682         COLNNOT16         GI 56805           1         SEQ ID NO:88         1298305         BRSTNOT07         GI 1019433           2         SEQ ID NO:89         1360501         LUNGNOT12         GI 2072705           3         SEQ ID NO:90         1362406         LUNGNOT12         GI 205250           4         SEQ ID NO:91         1405329         LATRTUT02         GI 205250           5         SEQ ID NO:92         1415523         BRAINOT12         GI 205250           6         SEQ ID NO:94         1418517         KIDNINOT09         GI 1065459           7         SEQ ID NO:95         1438165         PANCNOT08         GI 155161           8         SEQ ID NO:97         1510839         LUNGNOT14         GI 2145052           10         SEQ ID NO:99         1534876         SPLNNOT04         GI 496667           23         SEQ ID NO:99         1559131         SPLNNOT03         GI 206547           24         SEQ ID NO:100         1601473         BLADNOT03         GI 2196924           25         SEQ ID NO:102         1638407         UTRSNOT06         GI 2196024	SEO ID NO:8	SEQ ID NO:85	1273453	TESTTUT02		•
0         SEQ ID NO:87         1281682         COLNNOT16           1         SEQ ID NO:88         1298305         BRSTNOT07           2         SEQ ID NO:89         1360501         LUNGNOT12         GI 1019433           3         SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           4         SEQ ID NO:91         1405329         LATRTUT02         GI 205250           5         SEQ ID NO:92         1415523         BRAINOT12         GI 205250           6         SEQ ID NO:93         1416553         BRAINOT12         GI 205250           7         SEQ ID NO:94         1418517         KIDNNOT09         GI 1065459           8         SEQ ID NO:95         1440381         THYRNOT03         GI 1065459           9         SEQ ID NO:96         1440381         THYRNOT04         GI 2145052           10         SEQ ID NO:99         1559131         SPLNNOT04         GI 296667           22         SEQ ID NO:99         1559131         SPLNNOT04         GI 296667           23         SEQ ID NO:100         1601473         BLADNOT03         GI 2196924           24         SEQ ID NO:102         1634813         COLNNOT19         GI 2196924           25 <td>SEO ID NO:9</td> <td>SEQ ID NO:86</td> <td>1275261</td> <td>TESTTUT02</td> <td>GI 56805</td> <td>Rattus norvegicus</td>	SEO ID NO:9	SEQ ID NO:86	1275261	TESTTUT02	GI 56805	Rattus norvegicus
SEQ ID NO:88         1298305         BRSTNOT07           SEQ ID NO:89         1360501         LUNGNOT12         GI 1019433           SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           SEQ ID NO:91         1405329         LATRTUT02         GI 205250           SEQ ID NO:92         1416553         BRAINOT12         GI 205250           SEQ ID NO:93         1416553         BRAINOT12         GI 1065459           SEQ ID NO:94         1418517         KIDNNOT09         GI 1065459           SEQ ID NO:95         1440381         THYRNOT03         GI 1065459           SEQ ID NO:96         1534876         SPLNNOT04         GI 2145052           SEQ ID NO:99         1534876         SPLNNOT04         GI 496667           SEQ ID NO:99         1559131         SPLNNOT03         GI 496667           SEQ ID NO:100         1601473         BLADNOT03         GI 2196924           SEQ ID NO:101         1615809         BRAITUT12         GI 200547           SEQ ID NO:102         1634813         COLNNOT19         GI 2196924	SEO ID NO:10	SEQ ID NO:87	1281682	COLNNOT16		
SEQ ID NO:89         1360501         LUNGNOT12         GI 1019433           SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           SEQ ID NO:91         1405329         LATRTUT02         GI 205250           SEQ ID NO:92         1415223         BRAINOT12         GI 205250           SEQ ID NO:93         1416553         BRAINOT12         GI 205250           SEQ ID NO:94         1418517         KIDNNOT09         GI 1515161           SEQ ID NO:95         1440381         THYRNOT03         GI 1065459           SEQ ID NO:96         1440381         THYRNOT04         GI 2145052           SEQ ID NO:97         1510839         LUNGNOT14         GI 2145052           SEQ ID NO:99         1534876         SPLNNOT04         GI 296667           SEQ ID NO:100         1601473         BLADNOT03         GI 2196924           SEQ ID NO:101         1615809         BRAITUT12         GI 2196924           SEQ ID NO:102         1634813         COLNNOT19         GI 2196924	SEO ID NO:11	SEQ ID NO:88	1298305	BRSTNOT07		•
SEQ ID NO:90         1362406         LUNGNOT12         GI 2072705           SEQ ID NO:91         1405329         LATRTUT02         GI 205250           SEQ ID NO:92         1415223         BRAINOT12         GI 205250           SEQ ID NO:93         1416553         BRAINOT12         GI 205250           SEQ ID NO:94         1418517         KIDNNOT09         GI 1515161           SEQ ID NO:95         1440381         THYRNOT03         GI 1065459           SEQ ID NO:96         1440381         THYRNOT14         GI 2145052           SEQ ID NO:97         1510839         LUNGNOT14         GI 2145052           SEQ ID NO:99         1534876         SPLNNOT04         GI 496667           SEQ ID NO:100         1601473         BLADNOT03         GI 2196924           SEQ ID NO:101         1615809         BRAITUT12         GI 2196924           SEQ ID NO:102         1634813         COLNNOT19         GI 200547	SEO ID NO:12	SEO ID NO:89	1360501	LUNGNOT12	GI 1019433	Trypanosoma cruzi
SEQ ID NO:91       1405329       LATRTUT02         SEQ ID NO:92       1415223       BRAINOT12       GI 205250         SEQ ID NO:93       1416553       BRAINOT12         SEQ ID NO:94       1418517       KIDNNOT09       GI 1515161         SEQ ID NO:95       1438165       PANCNOT08       GI 1065459         SEQ ID NO:97       1510839       LUNGNOT14       GI 2145052         SEQ ID NO:99       1534876       SPLNNOT04       GI 496667         SEQ ID NO:99       1559131       SPLNNOT04       GI 496667         SEQ ID NO:100       1601473       BLADNOT03       GI 2196924         SEQ ID NO:101       1615809       BRAITUT12       GI 200547         SEQ ID NO:102       1634813       COLNNOT19       GI 200547	SEO ID NO:13	SEO ID NO:90	1362406	LUNGNOT12	GI 2072705	Mycobacterium tuberculosis
SEQ ID NO:92         1415223         BRAINOT12         GI 205250         J           SEQ ID NO:93         1416553         BRAINOT12         GI 205250         J           SEQ ID NO:94         1418517         KIDNNOT09         GI 1515161           SEQ ID NO:95         1440381         THYRNOT03         GI 1065459           SEQ ID NO:96         1440381         THYRNOT04         GI 2145052           SEQ ID NO:97         1510839         LUNGNOT14         GI 2145052           SEQ ID NO:99         1559131         SPLNNOT04         GI 496667           SEQ ID NO:100         1601473         BLADNOT03         GI 2196624           SEQ ID NO:101         1615809         BRAITUT12         GI 2196924           SEQ ID NO:102         1634813         COLNNOT19         GI 2196924           SEQ ID NO:103         1638407         UTRSNOT06         GI 200547	SEO ID NO:14	SEO ID NO:91	1405329	LATRTUT02		
SEQ ID NO:93       1416553       BRAINOT12         SEQ ID NO:94       1418517       KIDNNOT09         SEQ ID NO:95       1438165       PANCNOT08       GI 1515161         SEQ ID NO:96       1440381       THYRNOT03       GI 1065459         SEQ ID NO:97       1510839       LUNGNOT14       GI 2145052         SEQ ID NO:98       1534876       SPLNNOT04       GI 496667         SEQ ID NO:99       1559131       SPLNNOT04       GI 496667         SEQ ID NO:100       1601473       BLADNOT03       GI 2196924         SEQ ID NO:102       1634813       COLNNOT19       GI 2106924         SEQ ID NO:103       1638407       UTRSNOT06       GI 200547	SEO ID NO:15	SEQ ID NO:92	1415223	<b>BRAINOT12</b>	GI 205250	Rattus norvegicus
SEQ ID NO:94       1418517 KIDNNOT09         SEQ ID NO:95       1438165 PANCNOT08       GI 1515161         SEQ ID NO:96       1440381 THYRNOT03       GI 1065459         SEQ ID NO:97       1510839 LUNGNOT14       GI 2145052         SEQ ID NO:98       1534876 SPLNNOT04       GI 496667         SEQ ID NO:99       1559131 SPLNNOT04       GI 496667         SEQ ID NO:100       1601473 BLADNOT03       GI 2196924         SEQ ID NO:101       1615809 BRAITUT12       GI 2196924         SEQ ID NO:102       1634813 COLNNOT19       GI 2196924         SEQ ID NO:103       1638407 UTRSNOT06       GI 200547	SEO ID NO:16	SEQ ID NO:93	1416553	<b>BRAINOT12</b>		
SEQ ID NO:95       1438165       PANCNOT08       GI 1515161         SEQ ID NO:96       1440381       THYRNOT03       GI 1065459         SEQ ID NO:97       1510839       LUNGNOT14       GI 2145052         SEQ ID NO:98       1534876       SPLNNOT04       GI 496667         SEQ ID NO:99       1559131       SPLNNOT04       GI 496667         SEQ ID NO:100       1601473       BLADNOT03       SEQ ID NO:101         SEQ ID NO:101       1615809       BRAITUT12       GI 2196924         SEQ ID NO:102       1634813       COLNNOT19       GI 200547         SEQ ID NO:103       1638407       UTRSNOT06       GI 200547	SEO ID NO:17	SEQ ID NO:94	1418517	KIDNNOT09		
SEQ ID NO:96       1440381       THYRNOT03       GI 1065459         SEQ ID NO:97       1510839       LUNGNOT14       GI 2145052         SEQ ID NO:98       1534876       SPLNNOT04       GI 496667         SEQ ID NO:99       1559131       SPLNNOT04       GI 496667         SEQ ID NO:100       1601473       BLADNOT03       SEQ ID NO:101         SEQ ID NO:101       1615809       BRAITUT12       GI 2196924         SEQ ID NO:102       1634813       COLNNOT19       GI 200547         SEO ID NO:103       1638407       UTRSNOT06       GI 200547	SEQ ID NO:18	SEQ I	1438165	PANCNOT08	GI 1515161	Caenorhabditis elegans
SEQ ID NO:97       1510839 LUNGNOT14       GI 2145052         SEQ ID NO:98       1534876 SPLNNOT04       GI 496667         SEQ ID NO:99       1559131 SPLNNOT04       GI 496667         SEQ ID NO:100       1601473 BLADNOT03       SEQ ID NO:101         SEQ ID NO:101       1615809 BRAITUT12       SEQ ID NO:102         SEQ ID NO:102       1634813 COLNNOT19       GI 2196924         SEO ID NO:103       1638407 UTRSNOT06       GI 200547	SEQ ID NO:19		144038]	THYRNOT03	GI 1065459	Caenorhabditis elegans
SEQ ID NO:98       1534876       SPLNNOT04         SEQ ID NO:99       1559131       SPLNNOT04       GI 496667         SEQ ID NO:100       1601473       BLADNOT03         SEQ ID NO:101       1615809       BRAITUT12         SEQ ID NO:102       1634813       COLNNOT19       GI 2196924         SEQ ID NO:103       1638407       UTRSNOT06       GI 200547	<b>SEQ ID NO:20</b>	SEQ ID NO:97	1510839	LUNGNOT14	GI 2145052	Plasmodium bergnei
SEQ ID NO:99       1559131 SPLNNOT04       GI 496667         SEQ ID NO:100       1601473 BLADNOT03         SEQ ID NO:101       1615809 BRAITUT12         SEQ ID NO:102       1634813 COLNNOT19 GI 2196924         SEQ ID NO:103       1638407 UTRSNOT06 GI 200547	SEQ ID NO:21	SEQ ID NO:98	1534876	SPLNNOT04		
SEQ ID NO: 100       1601473       BLADNOT03         SEQ ID NO: 101       1615809       BRAITUT12         SEQ ID NO: 102       1634813       COLNNOT19       GI 2196924         SEQ ID NO: 103       1638407       UTRSNOT06       GI 200547	SEQ ID NO:22	SEQ ID NO:99	155913	SPLNNOT04	GI 496667	Saccharomyces cerevisiae
SEQ ID NO: 101         1615809         BRAITUT12           SEQ ID NO: 102         1634813         COLNNOT19         GI 2196924           SEQ ID NO: 103         1638407         UTRSNOT06         GI 200547	<b>SEQ ID NO:23</b>	SEQ ID NO:100		BLADNOT03		
SEQ ID NO: 102         1634813         COLNNOT19         GI 2196924           SEQ ID NO: 103         1638407         UTRSNOT06         GI 200547	SEO ID NO:24	SEQ ID NO:101	161580	BRAITUT12		
SEO ID NO: 103 1638407 UTRSNOT06 GI 200547	SEO ID NO:25	SEQ ID NO:102		COLNNOT19	GI 2196924	Mus musculus
7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7	<b>SEQ ID NO:26</b>	<b>SEQ ID NO:103</b>		7 UTRSNOT06	GI 200547	Mus musculus

## Hall fifth fifth the train that their seals at such stem and the fifth trains.

### $\Gamma ABI E$

Protein	Nucleotide	Clone ID Library	Library	NCBI I.D.	Homolog species
SEQ ID NO:27	SEQ ID NO:104	1653112	1653112 PROSTUT08	GI 49794	Mus musculus
SEQ ID NO:28	SEQ ID NO:105	1664634	1664634 BRSTNOT09	GI 1890375	Caenorhabditis elegans
<b>SEQ ID NO:29</b>	SEQ ID NO:106	1690990	1690990 PROSTUT10		
<b>SEQ ID NO:30</b>	<b>SEQ ID NO:107</b>	1704050	1704050 DUODNOT02	GI 1814277	Homo sapiens
<b>SEQ ID NO:31</b>	SEQ ID NO:108	1711840	1711840 PROSNOT16	GI 182651	Homo sapiens
<b>SEQ ID NO:32</b>	SEQ ID NO:109	1747327	1747327 STOMTUT02	GI 2062391	Homo sapiens
SEQ ID NO:33	SEQ ID NO:110	1750632	1750632 STOMTUT02	GI 459002	Caenorhabditis elegans
SEQ ID NO:34	SEQ ID NO:111	1812375	1812375 PROSTUT12		
<b>SEQ ID NO:35</b>	SEQ ID NO:112	1818761	1818761 PROSNOT20	GI 2493789	Homo sapiens
SEQ ID NO:36	SEQ ID NO:113	1824469	1824469 GBLATUT01	GI 2052134	Mycobacterium tuberculosis
SEQ ID NO:37	SEQ ID NO:114	1864292	1864292 PROSNOT19	GI 295671	Saccharomyces cerevisiae
SEQ ID NO:38	SEQ ID NO:115	1866437	1866437 THP1NOT01		
<b>SEQ ID NO:39</b>	SEQ ID NO:116	1871375	1871375 SKINBIT01		
SEQ ID NO:40	SEQ ID NO:117	1880830	1880830 LEUKNOT03	GI 1872521	Arabidopsis thaliana
<b>SEQ ID NO:41</b>	SEQ ID NO:118	1905325	1905325 OVARNOT07	GI 1754971	Homo sapiens
<b>SEQ ID NO:42</b>	SEQ ID NO:119	1919931	1919931 BRSTTUT01	GI 2104517	Homo sapiens
<b>SEQ ID NO:43</b>	<b>SEQ ID NO:120</b>	1969426	1969426 BRSTNOT04		
SEQ ID NO:44	SEQ ID NO:121	1969948	1969948 UCMCL5T01		
<b>SEQ ID NO:45</b>	SEQ ID NO:122	1988911	1988911 LUNGAST01	GI 56649	Rattus norvegicus
<b>SEQ ID NO:46</b>	SEQ ID NO:123	2061561	2061561 OVARNOT03		
<b>SEQ ID NO:47</b>	SEQ ID NO:124	2084489	2084489 PANCNOT04	GI 2262136	Arabidopsis thaliana
<b>SEQ ID NO:48</b>	SEQ ID NO:125	2203226	2203226 SPLNFET02	GI 1911776	Homo sapiens
<b>SEQ ID NO:49</b>	SEQ ID NO:126	2232884	2232884 PROSNOT16		
<b>SEQ ID NO:50</b>	SEQ ID NO:127	2328134	2328134 COLNNOT11	GI 1911776	Homo sapiens
<b>SEQ ID NO:51</b>	SEQ ID NO:128	2382718	2382718 ISLTNOT01	GI 1814277	Homo sapiens
<b>SEQ ID NO:52</b>	SEQ ID NO:129	2452208	2452208 ENDANOT01		

### TABLE 1

Protein	Nucleotide	Clone ID Library	Library	NCBI I.D.	Homolog species
<b>SEQ ID NO:53</b>	SEQ ID NO:130	2457825	2457825 ENDANOT01	GI 1418625	Caenorhabditis elegans
SEQ ID NO:54	SEQ ID NO:131	2470740	2470740 THP1NOT03		
<b>SEQ ID NO:55</b>	SEQ ID NO:132	2479092	2479092 SMCANOT01		
SEQ ID NO:56 SEQ II	SEQ ID NO:133	2480544	2480544 SMCANOT01	GI 169345	Phaseolus vulgaris
<b>SEQ ID NO:57</b>	SEQ ID NO:134	2518547	2518547 BRAITUT21	GI 33969	Homo sapiens
SEQ ID NO:58	SEQ ID NO:135	2530650	2530650 GBLANOT02	GI 2204111	Bos taurus
SEQ ID NO:59	SEQ ID NO:136	2652271	2652271 THYMNOT04	GI 895855	Solanum lycopersicum
SEQ ID NO:60	SEQ ID NO:137	2746976	2746976 LUNGTUT11	GI 191983	Mus musculus
SEQ ID NO:61	SEQ ID NO:138	2753496	2753496 THP1AZS08	GI 987286	Schizosaccharomyces pombe
<b>SEQ ID NO:62</b>	SEQ ID NO:139	2781553	2781553 OVARTUT03		
SEQ ID NO:63	SEQ ID NO:140	2821925	2821925 ADRETUT06		
SEQ ID NO:64	SEQ ID NO:141	2879068	2879068 UTRSTUT05	GI 870749	Homo sapiens
SEQ ID NO:65	SEQ ID NO:142	2886757	2886757 SINJNOT02	GI 1420026	Saccharomyces cerevisiae
SEQ ID NO:66	SEQ ID NO:143	2964329	2964329 SCORNOT04	GI 311667	Saccharomyces cerevisiae
SEQ ID NO:67	SEQ ID NO:144	2965248	2965248 SCORNOT04	GI 1478503	Homo sapiens
SEQ ID NO:68	SEQ ID NO:145	3000534	3000534 TLYMNOT06	GI 1741868	Homo sapiens
SEQ ID NO:69	SEQ ID NO:146	3046870	3046870 HEAANOT01	GI 1067079	Caenorhabditis elegans
SEQ ID NO:70	SEQ ID NO:147	3057669	3057669 PONSAZT01	GI 260241	
SEQ ID NO:71	SEQ ID NO:148	3088178	3088178 HEAONOT03	GI 498997	Saccharomyces cerevisiae
SEQ ID NO:72	SEQ ID NO:149	3094321	3094321 BRSTNOT19	GI 793879	Saccharomyces cerevisiae
SEQ ID NO:73	SEQ ID NO:150	3115936	3115936 LUNGTUT13	GI 517174	Saccharomyces cerevisiae
SEQ ID NO:74	SEQ ID NO:151	3116522	3116522 LUNGTUT13	GI 1669560	Homo sapiens
SEQ ID NO:75	SEQ ID NO:152	3117184	3117184 LUNGTUT13	GI 1418628	Caenorhabditis elegans
SEQ ID NO:76	SEQ ID NO:153	3125156	3125156 LNODNOT05	GI 804750	Homo sapiens
SEQ ID NO:77   SEQ I	SEQ ID NO:154	3129120	3129120 LUNGTUT12	GI 1256890	Saccharomyces cerevisiae

25

30

### PF-0459 US

Incyte Clone 305841 from the heart tissue cDNA library (HEARNOT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:78, was derived from Incyte Clones 305841 (HEARNOT01), 22049 (ADENINB01),168880 (LIVRNOT01), 1321915 (BLADNOT04), and the shotgun sequences SAWA02804, SAWA02781, SAWA01969, and SAWA01937.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:1. SIGP-1 is 348 amino acids in length and has a potential amidation site at Q120; a potential N-glycosylation site at N181; two potential casein kinase II phosphorylation sites at S19 and T279; a potential glycosaminoglycan attachment site at S35; and three potential protein kinase C phosphorylation sites at S19, S268, and S343. SIGP-1 shares 56% identity with human GP36b glycoprotein (GI 505652). The fragment of SEQ ID NO:78 including the 5' region from about nucleotide 117 to about nucleotide 161 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, neural, cardiovascular, hematopoietic and immune, and developmental cDNA libraries. Approximately 42% of these libraries are associated with neoplastic disorders, 28% with inflammation, and 21% with cell proliferation.

Nucleic acids encoding the SIGP-2 of the present invention were first identified in Incyte Clone 322866 from the eosinophil cDNA library (EOSIHET02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:79, was derived from Incyte Clones 322866 (EOSIHET02), 470107 (MMLR1DT01), 873933 (LUNGAST01), and 2268817 (UTRSNOT02)

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:2. SIGP-2 is 194 amino acids in length and has two potential N-glycosylation sites at N129 and N148; two potential casein kinase II phosphorylation sites at S74 and S151; four potential protein kinase C phosphorylation sites at S5, S74, S130, and S163; a potential tyrosine kinase phosphorylation site at Y171; two potential prokaryotic membrane lipoprotein lipid attachment sites at F15 and S61; and a transmembrane 4 protein family signature from G60 to L82. SIGP-2 shares 90% identity with CD53, a human cell surface antigen (GI 180141). The fragment of SEQ ID NO:79 from about nucleotide 624 to about nucleotide 686 is useful for hybridization. Northern analysis shows the expression of

30

5

10

### PF-0459 US

this sequence in hematopoietic and immune, gastrointestinal, cardiovascular, reproductive, musculoskeletal, and neural cDNA libraries. Approximately 54% of these libraries are associated with inflammation, 39% with neoplastic disorders, and 11% with cell proliferation.

Nucleic acids encoding the SIGP-3 of the present invention were first identified in Incyte Clone 546656 from the bronchial epithelium primary cell line cDNA library (BEPINOT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:80, was derived from Incyte Clones 546656 (BEPINOT01), 1316266 (BLADTUT02), 2095988 (BRAITUT02), 1318172 (BLADNOT04), 2809506 (TLYMNOT04), 1293412 and 1293630 (PGANNOT03), 2585048 (BRAITUT22), 2941370 (HEAONOT03), 2297230 (BRSTNOT05), 1233586 (LUNGFET03), and the shotgun sequence SAEA02986.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:3. SIGP-3 is 342 amino acids in length and has a potential amidation site at H4; a potential N-glycosylation site at N23; seven potential casein kinase II phosphorylation sites at S38, T90, T105, T124, S139, T284, and T324; three potential protein kinase C phosphorylation sites at S25, T71, and S200; two potential tyrosine kinase phosphorylation sites at Y13 and Y69; and a beta-transducin family Trp-Asp repeats signature sequence from I282 to I296. SIGP-3 shares 100% identity with human HAN11 (GI 2290530). The fragment of SEQ ID NO:80 from about nucleotide 107 to about nucleotide 139 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, cardiovascular, hematopoietic and immune, neural, urologic, and developmental cDNA libraries. Approximately 43% of these libraries are associated with neoplastic disorders, 25% with inflammation, and 20% with cell proliferation.

Nucleic acids encoding the SIGP-4 of the present invention were first identified in Incyte Clone 693453 from the synovial membrane cDNA library (SYNORAT03) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:81, was derived from Incyte Clones 693453 (SYNORAT03), 2505458 (CONUTUT01), 1527363 (UCMCL5T01), 1275308 (TESTTUT02), 1377126 (LUNGNOT10), 538256 (LNODNOT02), 3125441 (LNODNOT05), 1955296 (CONNNOT01), 1821536 (GBLATUT01), 2055631 (BEPINOT01), and 2028161 (KERANOT02).

25

30

5

10

## PF-0459 US

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:4. SIGP-4 is 656 amino acids in length and has a potential N-glycosylation site at N73; nine potential casein kinase II phosphorylation sites at S140, S191, T250, T252, S330, S340, S517, S617, and T630; a potential leucine zipper pattern from L430 to L451; four potential N-myristoylation sites at G77, G246, G484, and A651; eleven potential protein kinase C phosphorylation sites at S18, T90, S93, T318, S490, S503, S532, T565, T608, S609, and T629; and a potential tyrosine kinase phosphorylation site at Y326. SIGP-4 shares 20% identity with Caenorhabditis elegans protein encoded by T01G9.4 (GI 1419461). The fragment of SEQ ID NO:81 from about nucleotide 202 to about nucleotide 255 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, hematopoietic and immune, neural, and developmental cDNA libraries. Approximately 40% of these libraries are associated with neoplastic disorders, 30% with inflammation, and 30% with cell proliferation.

Nucleic acids encoding the SIGP-5 of the present invention were first identified in Incyte Clone 866885 from the brain tumor cDNA library (BRAITUT03) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:82, was derived from Incyte Clones 866885 (BRAITUT03), 2991983 (KIDNFET02), 067954 (HUVESTB01), and 1499109 (SINTBST01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:5. SIGP-5 is 236 amino acids in length and has a potential N-glycosylation site at N199; two potential casein kinase II phosphorylation sites at S8 and T72; a potential N-myristoylation site at G169; and three potential protein kinase C phosphorylation sites at T43, S96, and T201. SIGP-5 shares 24% identity with rat syntaxin (GI 1488683). The fragment of SEQ ID NO:82 from about nucleotide 43 to about nucleotide 93 is useful for hybridization. Northern analysis shows the expression of this sequence in hematopoietic and immune, reproductive, gastrointestinal, neural, cardiovascular, and developmental cDNA libraries. Approximately 43% of these libraries are associated with neoplastic disorders, 26% with inflammation, and 19% with cell proliferation.

Nucleic acids encoding the SIGP-6 of the present invention were first identified in Incyte Clone 1242271 from the lung tissue cDNA library (LUNGNOT03) using a computer

25

30

5

# PF-0459 US

search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:83, was derived from Incyte Clones 1242271 (LUNGNOT03), 968114 (BRSTNOT05), 1251728 (LUNGFET03), and the shotgun sequence SAZA00142.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:6. SIGP-6 is 195 amino acids in length and has a potential cAMP- and cGMP-dependent protein kinase phosphorylation site at S79; six potential casein kinase II phosphorylation sites at S79, T85, S113, T166, T171, and T188; three potential protein kinase C phosphorylation sites at S20, S150, and S185; and a potential mitochondrial energy transfer proteins signature from P25 to Y33. The fragment of SEQ ID NO:83 from about nucleotide 98 to about nucleotide 133 is useful for hybridization. Northern analysis shows the expression of this sequence in urologic, neural, reproductive, and cardiovascular cDNA libraries. Approximately 50% of these libraries are associated with neoplastic disorders, 14% with inflammation, and 21% with cell proliferation.

Nucleic acids encoding the SIGP-7 of the present invention were first identified in Incyte Clone 1255027 from the fetal lung cDNA library (LUNGFET03) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:84, was derived from Incyte Clones 1255027 (LUNGFET03), 2055704 (BEPINOT01), 1351096 (LATRTUT02), 835188 (PROSNOT07), and 1695810 (COLNNOT23).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:7. SIGP-7 is 608 amino acids in length and has a potential amidation site at T112; five potential N-glycosylation sites at N73, N110, N410, N436, and N478; two potential cAMP- and cGMP-dependent protein kinase phosphorylation sites at S123 and S185; ten potential casein kinase II phosphorylation sites at T2, S75, S166, S170, S185, S274, S463, S505, S517, and T588; and thirteen potential protein kinase C phosphorylation sites at T19, S32, S46, T112, T221, S274, S299, T337, S373, S412, S431, S438, and S555. SIGP-7 shares 16% identity with canine pinin (GI 1684845). The fragment of SEQ ID NO:84 from about nucleotide 181 to about nucleotide 219 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, neural, cardiovascular, and developmental cDNA libraries. Approximately 43% of these libraries are associated with neoplastic disorders, 21% with inflammation, and

30

5

10

PF-0459 US

20% with cell proliferation.

Nucleic acids encoding the SIGP-8 of the present invention were first identified in Incyte Clone 1273453 from the testicle cDNA library (TESTTUT02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:85, was derived from Incyte Clones 1273453 (TESTTUT02), 1970337 (UCMCL5T01), 1218926 (NEUTGMT01), 1881349 (LEUKNOT03), and 1722377 (BLADNT06).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:8. SIGP-8 is 267 amino acids in length and has a potential N glycosylation site at N230, five potential casein kinase II phosphorylation sites at S9, T45, T77, S190, and T263, and two potential protein kinase C phosphorylation sites at S232 and S236. The fragment of SEQ ID NO:85 from about nucleotide 140 to about nucleotide 175 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, cardiovascular, and hematopoietic and immune cDNA libraries. Approximately 42% of these libraries are associated with neoplastic disorders and 40% with immune response.

Nucleic acids encoding the SIGP-9 of the present invention were first identified in Incyte Clone 1275261 from the testicle cDNA library (TESTTUT02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:86, was derived from Incyte Clones 1275261 (TESTTUT02), 775078 (COLNNOT05), 514772 (MMLR1DT01), and 3224071 (COLNNON03).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:9. SIGP-9 is 285 amino acids in length and has a potential amidation site at S260, three potential N glycosylation sites at N85, N100 and N156, a potential cAMP- and cGMP-dependent protein kinase phosphorylation site at T168, three potential casein kinase II phosphorylation sites at T168, T215, and S230, three potential protein kinase C phosphorylation sites at S163, S230, and S260, and a potential tyrosine kinase phosphorylation site at Y72. SIGP-9 shares 24% identity with rat OX-45 antigen preprotein (GI 56805). The fragment of SEQ ID NO:86 from about nucleotide 243 to about nucleotide 293 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, and hematopoietic and immune cDNA

Am Am

25

30

5

10

# PF-0459 US

libraries. Approximately 50% of these libraries are associated with neoplastic disorders and 50% with immune response.

Nucleic acids encoding the SIGP-10 of the present invention were first identified in Incyte Clone 1281682 from the colon cDNA library (COLNNOT16) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:87, was derived from Incyte Clones 2681940 (SINIUCT01), 1335652 (COLNNOT13), 2079572 (UTRSNOT08), 627405 (PGANNOT01) and 1281682 and 1282887 (COLNNOT16).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:10. SIGP-10 comprises a peptide of 76 amino acids in length, and has a potential signal peptide sequence from M1 to S18. The fragment of SEQ ID NO:87 encoding the potential signal peptide sequence from about nucleotide 908 through 970 is useful for hybridization. Northern analysis shows the expression of this sequence in gastrointestinal, neural, reproductive, and hematopoietic and immune cDNA libraries. Approximately 32% of these libraries are associated with neoplastic disorders and 53% with immune response.

Nucleic acids encoding the SIGP-11 of the present invention were first identified in Incyte Clone 1298305 from the breast cDNA library (BRSTNOT09) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:88, was derived from Incyte Clones 1298305 (BRSTNOT09), 3451203 (UTRSNON03), 2529672 (GBLAN0502), 2780863 (OVARTUT03), 927988 (BRAINOT04), 1684424 (PROSNOT15), 2243053 (PANCTUT02), and shotgun sequences SANA03310 and SANA00700.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:11. SIGP-11 is 147 amino acids in length and has a prokaryotic membrane lipoprotein lipid attachment site from L34 through C44. SIGP-11 also has a potential cAMP- and cGMP-dependent protein kinase phosphorylation site at S91, and a potential protein kinase C phosphorylation site at S13. The fragment of SEQ ID NO:88 from about nucleotide 1561 to about nucleotide 1611 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, and neural cDNA libraries. Approximately 50% of these libraries are associated with

neoplastic disorders and 22% with immune response.

Nucleic acids encoding the SIGP-12 of the present invention were first identified in Incyte Clone 1360501 from the lung cDNA library (LUNGNOT12) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:89, was derived from Incyte Clones 1360501 (LUNGNOT12), 2121661 (BRSTNOT07), 1706518 (DUODNOT02) and shotgun sequences SAJA02519, SAJA00749, SAJA01160, and SANA00513.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:12. SIGP-12 is 261 amino acids in length and has six potential N glycosylation sites at N19, N28, N98, N104, N164 and N178. SIGP-12 also has five potential casein kinase II phosphorylation sites at T82, S83, T91, T160, and S233, and nine potential protein kinase C phosphorylation sites at T35, T60, T82, S121, S131, T184, S233, S237, and T242. SIGP-12 shares 22% identity with <a href="Trypanosoma cruzi">Trypanosoma cruzi</a> mucin-like protein (GI 1019433). In addition, SIGP-12 shares two potential phosphorylation sites and a potential N-glycosylation site with the mucin-like protein. The fragment of SEQ ID NO:89 from about nucleotide 183 to about nucleotide 236 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, cardiovascular, and gastrointestinal cDNA libraries. Approximately 39% of these libraries are associated with neoplastic disorders and 26% with immune response.

Nucleic acids encoding the SIGP-13 of the present invention were first identified in Incyte Clone 1362406 from the lung cDNA library (LUNGNOT12) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:90, was derived from Incyte Clones 1362406 (LUNGNOT12), 1854401 (HNT3AZT01), 1570003 (UTRSNOT05) and shotgun sequences SANA03704, SANA00366, and SANA02152.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:13. SIGP-13 is 213 amino acids in length and has three potential protein kinase C phosphorylation sites at T40, S136, and T166. In addition, SIGP-13 has a highly hydrophobic signal peptide sequence from residue M1 to E34. SIGP-13 shares 20% identity with a Mycobacterium tuberculosis membrane protein (GI 2072705). The fragment of SEQ ID NO:90 encoding the potential signal peptide sequence

20

25

30

5

10

10 and and an in the second of the second of

25

30

5

domain from about nucleotide 157 to about nucleotide 219 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, developmental, neural, and cardiovascular cDNA libraries. Approximately 50% of these libraries are associated with neoplastic disorders and 18% with immune response.

Nucleic acids encoding the SIGP-14 of the present invention were first identified in Incyte Clone 1405329 from the heart cDNA library (LATRTUT02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:91, was derived from Incyte Clones 1405329 (LATRTUT02), and 2830813 (TLYMNOT03).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:14. SIGP-14 is 67 amino acids in length and has a cell attachment sequence comprising R13 through D15. In addition, SIGP-14 has a potential casein kinase II phosphorylation site at T12, and a potential protein kinase C phosphorylation site at T42. The fragment of SEQ ID NO:91 from about nucleotide 36 to about nucleotide 95 is useful for hybridization. Northern analysis shows the expression of this sequence in cardiovascular, developmental, reproductive, and hematopoietic and immune cDNA libraries. Approximately 43% of these libraries are associated with neoplastic disorders and 21% with immune response.

Nucleic acids encoding the SIGP-15 of the present invention were first identified in Incyte Clone 1415223 from the brain cDNA library (BRAINOT12) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:92, was derived from Incyte Clones 1415223 (BRAINOT12) and 529786 (BRAINOT03).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:15. SIGP-15 is 161 amino acids in length and has a potential N-glycosylation site at N57, two potential casein kinase II phosphorylation sites at S84 and S96, and five potential protein kinase C phosphorylation sites at S11, T62, S75, S83, and S84. SIGP-15 shares 30% identity with rat Ly6C antigen (GI 205250). The fragment of SEQ ID NO:92 from about nucleotide 28 to about nucleotide 81 is useful for hybridization. Northern analysis shows the expression of this sequence in developmental, reproductive, and neural cDNA libraries. Approximately 33% of these libraries are associated with neoplastic disorders, 33% with cell proliferation, and 17% with immune response.

The state of the s

5

10

Nucleic acids encoding the SIGP-16 of the present invention were first identified in Incyte Clone 1416553 from the brain cDNA library (BRAINOT12) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:93, was derived from Incyte Clones 1416553 (BRAINOT12), 663124 (BRAINOT03) and shotgun sequences SANA01409, SANA03513, and SANA02713.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:16. SIGP-16 is 141 amino acids in length and has a glycosaminoglycan attachment site at S20. In addition, SIGP-16 has a potential casein kinase II phosphorylation site at S61, and a potential protein kinase C phosphorylation site at S53. The fragment of SEQ ID NO:93 from about nucleotide 784 to about nucleotide 831 is useful for hybridization. Northern analysis shows the expression of this sequence in neural cDNA libraries. Approximately 27% of these libraries are associated with neoplastic disorders, and 27% with neurological disorders.

Nucleic acids encoding the SIGP-17 of the present invention were first identified in Incyte Clone 1418517 from the kidney cDNA library (KIDNNOT09) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:94, was derived from Incyte Clones 1418517 (KIDNNOT09), 2456866 (ENDANOT01), 136927 (SYNORAB01), 1620442 (BRAITUT13), 1492394 (PROSNON01), 1534435 (SPLNNOT04), and 2505923 (CONUTUT01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:17. SIGP-17 is 152 amino acids in length and has a potential N glycosylation site at N76; a potential cAMP- and cGMP-dependent protein kinase phosphorylation site at T67; four potential casein kinase II phosphorylation sites at S9, T30, S107, and S124; and three potential protein kinase C phosphorylation sites at T30, S34, and T78. The fragment of SEQ ID NO:94 from about nucleotide 49 to about nucleotide 99 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, cardiovascular, musculoskeletal, and gastrointestinal cDNA libraries. Approximately 44% of these libraries are associated with neoplastic disorders, 23% with immune response, and 20% with cell proliferation.

Nucleic acids encoding the SIGP-18 of the present invention were first identified in

25

30

5

10

### PF-0459 US

Incyte Clone 1438165 from the pancreas cDNA library (PANCNOT08) using a computer search for amino acid alignments. A consensus sequence, SEQ ID NO:95, was derived from Incyte Clones 360389 (SYNORAB01), 485693 (HNT2RAT01), 1233177 (LUNGFET03), 1255551 (MENITUT03),1438165 (PANCNOT08),1554990 (BLADTUT04), and shotgun sequences SAOA00854 and SAOA00855.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:18. SIGP-18 is 742 amino acids in length and has a potential N-glycosylation site at N448; a microbodies C-terminal targeting signal in the triplet N740HL; twelve potential casein kinase II phosphorylation sites at S3, S53, S120, T122, T169, T178, S179, S195, T284, S290, S400, and S573; five potential protein kinase C phosphorylation sites at T178, S195, S208, S299, and S364; and two potential tyrosine kinase phosphorylation sites at Y296 and Y512. Cysteine residues, representing potential intramolecular disulfide bridging sites, are found at residues C87, C204, C312, C339, C343, C469, C497, C558, C657, C693, and C720. SIGP-18 shares 19% homology with C. elegans protein encoded by M163.4 (GI 1515161), including eight of the eleven cysteine residues found in SIGP-18. The fragment of SEQ ID NO:95 from about nucleotide 322 to about nucleotide 387 is useful for hybridization. Northern analysis shows the expression of this sequence in cardiovascular, male and female reproductive, and gastrointestinal cDNA libraries. Approximately 44% of these libraries are associated with neoplastic disorders, 23% with inflammation and the immune response, and 19% with fetal development.

Nucleic acids encoding the SIGP-19 of the present invention were first identified in Incyte Clone 1440381 from the thyroid cDNA library (THYRNOT03) using a computer search for amino acid alignments. A consensus sequence, SEQ ID NO:96, was derived from Incyte Clones 989671 (COLNNOT11),1440381 (THYRNOT03), 3507668 (CONCNOT01), and shotgun sequences SAOA03364, SAOA02692, SAOA00489, SAOA02355, SAOA02405, SAOA01209, SAOA00809, and SAOA00274.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:19. SIGP-19 is 805 amino acids in length and has three potential N-glycosylation sites at N211, N215, and N327; one cAMP- and cGMP-dependent protein kinase potential phosphorylation sites at T749; sixteen potential casein kinase II

30

5

10

phosphorylation sites at S8, T54, T175, T228, S229, S250, S292, S329, T390, S401, S415, S471, S492, S671, T780, and S795; ten potential protein kinase C phosphorylation sites at S206, T396, S401, S442, T455, S600, S671, T683, S730, and S795; and two potential tyrosine kinase phosphorylation sites at Y437 and Y476. SIGP-19 shares 33% homology with a ubiquitin-conjugating, E2-like enzyme from C. elegans (GI 1065459). Both molecules share a "UBC domain" characteristic of ubiquitin-conjugating enzymes extending from approximately residue V559 to I647 of SIGP-19, and containing an active site cysteine residue, C614, required for thiolester formation. A characteristic proline-rich region, found at the N-terminal end of the UBC domain and extending from approximately P564 to P589 in SIGP-19, is also shared by both proteins. The fragment of SEQ ID NO:96 from about nucleotide 1678 to about nucleotide 1800 is useful for hybridization. Northern analysis shows the expression of this sequence in cardiovascular and male and female reproductive cDNA libraries. Approximately 50% of these libraries are associated with neoplastic disorders, 14% with inflammation and the immune response, and 19% with fetal development.

Nucleic acids encoding the SIGP-20 of the present invention were first identified in Incyte Clone 1510839 from the lung cDNA library (LUNGNOT14) using a computer search for amino acid alignments. A consensus sequence, SEQ ID NO:97, was derived from Incyte Clones 962326 (BRSTTUT03), 1383254 (BRAITUT08), 1510839 (LUNGNOT14), 1970949 (UCMCL5T01), 2214224 (SINTFET03), and shotgun sequences SAOA01059 and SAOA02595.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:20. SIGP-20 is 195 amino acids in length and has a potential signal peptide sequence between M1 and A39. SIGP-20 also has a potential N-glycosylation site at N83; and three potential casein kinase II phosphorylation sites at T161, T169, and T181; and three potential protein kinase C phosphorylation sites at T121, T143, and T153. SIGP-20 shares 21% homology with <u>Plasmodium berghei</u> merozoite surface protein-1 (GI 2145052). The fragment of SEQ ID NO:97 from about nucleotide 439 to about nucleotide 502 is useful for hybridization. Northern analysis shows the expression of this sequence in cardiovascular, male and female reproductive, and developmental cDNA libraries.

30

5

10

## PF-0459 US

Approximately 48% of these libraries are associated with neoplastic disorders, 13% with inflammation and the immune response, and 19% with fetal development.

Nucleic acids encoding the SIGP-21 of the present invention were first identified in Incyte Clone 1534876 from the spleen cDNA library (SPLNNOT04) using a computer search for amino acid alignments. A consensus sequence, SEQ ID NO:98, was derived from Incyte Clones 1253004 (LUNGFET03), 1382838 (BRAITUT08), 1532501 (SPLNNOT04), 1534876 (SPLNNOT04), 1705806 (DUODNOT02), 1738301 (COLNNOT22), 1926209 (BRSTNOT02), and shotgun sequences SAOA00587, SAOA02048, and SAOA03535.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:21. SIGP-21 is 161 amino acids in length and has a potential signal peptide sequence between M1 and C13. SIGP-21 also has 17 cysteine residues with the potential for forming intramolecular disulfide bridges. Six of these cysteine residues, between residues C129 and C152, are found in a signature sequence for trypsin/alpha-amylase inhibitors that form a structure with intramolecular disulfide bridges. SIGP-21 has two potential casein kinase II phosphorylation sites at T25 and S35; and two potential protein kinase C phosphorylation sites at S35 and T87. The fragment of SEQ ID NO:98 from about nucleotide 406 to about nucleotide 477, which encompasses the trypsin/alpha-amylase inhibitor signature sequence, is useful for hybridization. Northern analysis shows the expression of this sequence in gastrointestinal and male and female reproductive cDNA libraries. Approximately 45% of these libraries are associated with neoplastic disorders and 28% with inflammation and the immune response.

Nucleic acids encoding the SIGP-22 of the present invention were first identified in Incyte Clone 1559131 from the spleen cDNA library (SPLNNOT04) using a computer search for amino acid alignments. A consensus sequence, SEQ ID NO:99, was derived from Incyte Clones 1559131 (SPLNNOT04), 1671080 (BMARNOT03), 1924001 (BRSTTUT01), and shotgun sequences SAPA01073 and SAOA02895.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:22. SIGP-22 is 160 amino acids in length and has cysteine residues capable of forming intramolecular disulfide bridges at C40, C47, C108, C114, C129, C154, and C158. SIGP-22 has one potential casein kinase II phosphorylation site at S9 and

30

5

10

## PF-0459 US

one potential protein kinase C phosphorylation site at S31. SIGP-22 shares 26% homology with C-215 protein from Saccharomyces cerevisiae (GI 496667), including four of the cysteine residues found in SIGP-22. The fragment of SEQ ID NO:99 from about nucleotide 154 to about nucleotide 193 is useful for hybridization. Northern analysis shows the expression of this sequence in hematopoietic and male and female reproductive cDNA libraries. Approximately 33% of these libraries are associated with neoplastic disorders and 67% with the immune response.

Nucleic acids encoding the SIGP-23 of the present invention were first identified in Incyte Clone 1601473 from the bladder cDNA library (BLADNOT03) using a computer search for amino acid alignments. A consensus sequence, SEQ ID NO:100, was derived from Incyte Clones 1601473 (BLADNOT03), and shotgun sequences SAOA00407, SAOA02497, SAOA02747, and SAOA02958.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:23. SIGP-23 is 76 amino acids in length and has two cysteine residues with the potential of forming an intramolecular disulfide bridge at C58 and C72. SIGP-23 has one potential casein kinase II phosphorylation site at S7 and three potential protein kinase C phosphorylation sites at S7, T29, and T46. The fragment of SEQ ID NO:100 from about nucleotide 139 to about nucleotide 180 is useful for hybridization. Northern analysis shows the expression of this sequence in breast, brain, spleen, thyroid, and bladder cDNA libraries. Approximately 33% of these libraries are associated with neoplastic disorders, 17% with neural disorders, and 17% with immune disorders.

Nucleic acids encoding the SIGP-24 of the present invention were first identified in Incyte Clone 1615809 from the brain tumor cDNA library (BRAITUT12) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:101, was derived from Incyte Clones 1615809 (BRAITUT12), 924499 (BRAINOT04), 1273065 (TESTTUT02), 1517058 (PANCTUT01), 1596867 (BRAINOT14), and 1361446 (LUNGNOT12), and shotgun sequence SAOA02975.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:24. SIGP-24 is 336 amino acids in length and has 13 potential phosphorylation sites at T27, T72, S74, S76, T99, S104, S109, S140, S178, S210, T281,

30

5

10

PF-0459 US

S326, S39. SIGP-24 also has a potential signal peptide sequence between M1 and Y18. The fragment of SEQ ID NO:101 from about nucleotide 187 to about nucleotide 247 is useful for hybridization. Northern analysis shows the expression of this sequence in cardiovascular, gastrointestinal, neural, and reproductive cDNA libraries. Approximately 48% of these libraries are associated with neoplastic disorders and 21% with immune response.

Nucleic acids encoding the SIGP-25 of the present invention were first identified in Incyte Clone 1634813 from the cecal tissue cDNA library (COLNNOT19) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:102, was derived from Incyte Clones 1634813 (COLNNOT19), 2904583 (THYMNOT05), 1634813 (COLNNOT19), and 1310492 (COLNFET02), and shotgun sequence SAPA04436.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:25. SIGP-25 is 150 amino acids in length and has one potential N-glycosylation site at N139; and five potential phosphorylation sites at T48, S118, S126, S135, and S136. SIGP-25 also has a potential signal peptide sequence encompassing residues M1-A23. SIGP-25 shares 28% identity with mouse beta chemokine, Exodus-2 (GI 2196924). The fragment of SEQ ID NO:102 from about nucleotide 175 to about nucleotide 235 is useful for hybridization. Northern analysis shows the expression of this sequence in gastrointestinal, developmental, hematopoietic, and immunological cDNA libraries. Approximately 50% of these libraries are associated with fetal development/cell proliferation and 25% with immune response.

Nucleic acids encoding the SIGP-26 of the present invention were first identified in Incyte Clone 1638407 from the myometrial tissue cDNA library (UTRSNOT06) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:103, was derived from Incyte Clones 1638407 (UTRSNOT06), 3541410 (SEMVNOT04), 1290413 (BRAINOT11), 1467841 (PANCTUT02), 1306495 (PLACNOT02), and 1907983 (CONNTUT01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:26. SIGP-26 is 217 amino acids in length and has seven potential phosphorylation sites at T214, S68, S148, S189, S30, S110, and Y149. SIGP-26 also has a potential signal peptide sequence between M1 and G31. SIGP-26 shares 18%

30

5

10

## PF-0459 US

identity with a mouse proline-rich protein (GI 200547). The fragment of SEQ ID NO:103 from about nucleotide 146 to about nucleotide 206 is useful for hybridization. Northern analysis shows the expression of this sequence in gastrointestinal, hematopoietic, immunological, and reproductive cDNA libraries. Approximately 42% of these libraries are associated with neoplastic disorders and 39% with immune response.

Nucleic acids encoding the SIGP-27 of the present invention were first identified in Incyte Clone 1653112 from the prostate tumor tissue cDNA library (PROSTUT08) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:104, was derived from Incyte Clones 1653112 (PROSTUT08), 3450102 (UTRSNON03), 1969850 (UCMCL5T01), 1880259 (LEUKNOT03), 1504393 (BRAITUT07), and 394029 (TMLR2DT01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:27. SIGP-27 is 504 amino acids in length and has eight potential phosphorylation sites at T338, T13, S38, T56, T132, T490, S33, and T472. SIGP-27 also has one potential leucine zipper pattern between L418 and L439. SIGP-27 shares 16% identity with mouse alpha-1 type-X collagen (GI 49794). The fragment of SEQ ID NO:104 from about nucleotide 130 to about nucleotide 190 is useful for hybridization. Northern analysis shows the expression of this sequence in cardiovascular, endocrine, hematopoietic, immunological, neural, and reproductive cDNA libraries. Approximately 55% of these libraries are associated with neoplastic disorders and 22% with immune response.

Nucleic acids encoding the SIGP-28 of the present invention were first identified in Incyte Clone 1664634 from the breast tissue cDNA library (BRSTNOT09) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:105, was derived from Incyte Clones 1664634 (BRSTNOT09) and 571656 (OVARNON01), and shotgun sequences SAPA04612, SAPA00377, and SAPA03034.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:28. SIGP-28 is 320 amino acids in length and has two potential N-glycosylation sites at N122 and N139; and eight potential phosphorylation sites at T30, S52, S109, S162, S220, S96, T258, and S280. SIGP-28 also has a potential signal peptide

30

5

10

sequence between M1 and A21. SIGP-28 shares 28% identity with a <u>C. elegans</u> protein encoded by F32A7.4 (GI 1890375). The fragment of SEQ ID NO:105 from about nucleotide 280 to about nucleotide 340 is useful for hybridization. Northern analysis shows the expression of this sequence in cardiovascular, gastrointestinal, hematopoietic, immunological, neural, and reproductive cDNA libraries. Approximately 38% of these libraries are associated with neoplastic disorders and 32% with immune response.

Nucleic acids encoding the SIGP-29 of the present invention were first identified in Incyte Clone 1690990 from the prostatic tumor tissue cDNA library (PROSTUT10) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:106, was derived from Incyte Clone 1690990 (PROSTUT10), and shotgun sequences SAPA01051, SAPA04063, SAPA01670, SAPA02170, SAPA01946, and SAPA00282.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:29. SIGP-29 is 117 amino acids in length and has one potential N-glycosylation site at N96; four potential phosphorylation sites at S16, S34, T78, and S62; and one potential N-myristoylation site at G5. SIGP-29 also has one potential microbodies C-terminal targeting signal at S115. The fragment of SEQ ID NO:106 from about nucleotide 1000 to about nucleotide 1062 is useful for hybridization. Northern analysis shows the expression of this sequence in gastrointestinal, reproductive, dermal, musculoskeletal, neural, and urogenital cDNA libraries. Approximately 77% of these libraries are associated with neoplastic disorders and 8% with immune response.

Nucleic acids encoding the SIGP-30 of the present invention were first identified in Incyte Clone 1704050 from the duodenal cDNA library (DUODNOT02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:107, was derived from Incyte Clones 865233 (BRAITUT03), 1359660 (LUNGNOT12), and 1704050 (DUODNOT02) and shotgun sequence SAPA02672.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:30. SIGP-30 is 298 amino acids in length and has one potential amidation site at P226; four potential N-glycosylation sites at N98, N187, N236, and N277; seven potential casein kinase II phosphorylation sites at T39, S59, T100, T149, S205, T284, and S286; three potential protein kinase C phosphorylation sites at T52, S58,

30

5

10

and S279; a potential signal sequence from M1 to G22; and a potential transmembrane spanning region from M230 to A261. SIGP-30 contains two potential immunoglobulin superfamily domains, from about F29 to about L131 and from about S138 to about R224. SIGP-30 shares 25% identity with the human A33 antigen precursor expressed in normal human colonic and small bowel epithelium and in human colon cancers (GI 1814277). In addition, the position of the hydrophobic transmembrane domain is conserved between these molecules. The cysteine residues at C50, C109, C139, C155, C214, and C254 are conserved between these molecules. The fragment of SEQ ID NO:107 from about nucleotide 1150 to about nucleotide 1209 is useful for hybridization. Northern analysis shows the expression of this sequence in neural, reproductive, cardiovascular, and endocrine cDNA libraries. Approximately 68% of these libraries are associated with cancer and 9% with immune response.

Nucleic acids encoding the SIGP-31 of the present invention were first identified in Incyte Clone 1711840 from the prostate cDNA library (PROSNOT16) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:108, was derived from Incyte Clones 1711840 (PROSNOT16) and 2550483 (LUNGTUT06) and shotgun sequence SAQA03185.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:31. SIGP-31 is 118 amino acids in length and has three potential protein kinase C phosphorylation sites at S48, T103, and S109; and a potential signal peptide sequence from M1 to A20. SIGP-31 shares 61% identity with human midkine, a retinoic acid-responsive heparin binding factor involved in regulation of growth and differentiation (GI 182651). The fragment of SEQ ID NO:108 from about nucleotide 511 to about nucleotide 555 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, developmental, neural, and cardiovascular cDNA libraries. Approximately 58% of these libraries are associated with cancer, 16% with immune response, and 23% with fetal/proliferating cells.

Nucleic acids encoding the SIGP-32 of the present invention were first identified in Incyte Clone 1747327 from the stomach tumor cDNA library (STOMTUT02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID

## PF-0459 US

NO:109, was derived from Incyte Clones 475228 (MMLR2DT01), 1500771 (SINTBST01), 1880656 (LEUKNOT03), 1747327 (STOMTUT02), and 2720285 (LUNGTUT10).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:32. SIGP-32 is 248 amino acids in length and has one potential N-glycosylation site at N56; three potential casein kinase II phosphorylation sites at S46, S134, and S140; and one potential protein kinase C phosphorylation site at T217. SIGP-32 shares 100% identity with human K12 protein precursor which is expressed in breast cancer cells and peripheral blood leukocytes (GI 2062391). Northern analysis shows the expression of this sequence in gastrointestinal, reproductive, hematopoietic/immune, and cardiovascular cDNA libraries. Approximately 59% of these libraries are associated with cancer and 35% with immune response.

Nucleic acids encoding the SIGP-33 of the present invention were first identified in Incyte Clone 1750632 from the stomach tumor cDNA library (STOMTUT02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:110, was derived from Incyte Clones 1521122 (BLADTUT04) and 1750632 (STOMTUT02) and shotgun sequences SAEA02182 and SAEA10021.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:33. SIGP-33 is 150 amino acids in length and has one potential protein kinase C phosphorylation site at S6. SIGP-33 shares 49% identity with the <u>C. elegans</u> protein encoded by R151.6 (GI 459002). The fragment of SEQ ID NO:110 from about nucleotide 514 to about nucleotide 573 is useful for hybridization. Northern analysis shows the expression of this sequence in cardiovascular and gastrointestinal cDNA libraries. Approximately 88% of these libraries are associated with cancer and 13% with immune response.

Nucleic acids encoding the SIGP-34 of the present invention were first identified in Incyte Clone 1812375 from the prostate tumor cDNA library (PROSTUT12) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:111, was derived from Incyte Clones 775001 (COLNNOT05), 834305 (PROSNOT07), 1504623 (BRAITUT07), and 1812375 (PROSTUT12) and shotgun sequences SAQA02414.

20

25

5

10

į

20

25

30

**#** 10

5

PF-0459 US

SATA00657, and SATA01478.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:34. SIGP-34 is 431 amino acids in length and has four potential N-glycosylation sites at N11, N49, N73, and N312; one potential cAMP- and cGMP-dependent protein kinase phosphorylation site at S197; six potential casein kinase II phosphorylation sites at T38, S79, S130, S165, S177, and T188; three potential protein kinase C phosphorylation sites at S184, T254, and S337; and a potential high affinity calcium ion-binding, vitamin K-dependent carboxylation domain between W371 and W408. The fragments of SEQ ID NO:111 from about nucleotide 222 to about nucleotide 282 and the potential carboxylation domain encoded from about nucleotide 1267 to about nucleotide 1380 are useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, neural, gastrointestinal, cardiovascular, and hematopoietic/immune DNA libraries. Approximately 52% of these libraries are associated with cancer, 24% with immune response, and 20% with fetal/proliferating cells.

Nucleic acids encoding the SIGP-35 of the present invention were first identified in Incyte Clone 1818761 from the prostate cDNA library (PROSNOT20) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:112, was derived from Incyte Clone 1818761 (PROSNOT20) and shotgun sequences SAJA00040, SAJA00601, SAJA01791, and SAJA02873.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:35. SIGP-35 is 278 amino acids in length and has one potential N-glycosylation site at N91; three potential casein kinase II phosphorylation sites at S9, S125, and S156; two potential protein kinase C phosphorylation sites at S77 and S224; one potential tyrosine kinase phosphorylation site at Y258; and a potential signal sequence from M1 to A30. SIGP-35 has fourteen consecutive collagen repeats (G-X-P or G-X-X) from G97 to P138 which could form a triple helical structure. SIGP-35 shares 28% identity with the human adipocyte complement-related protein precursor (Acrp30) (GI 2493789). The fragment of SEQ ID NO:112 from about nucleotide 157 to about nucleotide 210 is useful for hybridization. Northern analysis shows the expression of this sequence in developmental, dermal, gastrointestinal, hematopoietic/immune, neural, and

## PF-0459 US

reproductive cDNA libraries. Approximately 29% of these libraries are associated with cancer, 43% with immune response, and 29% with fetal development.

Nucleic acids encoding the SIGP-36 of the present invention were first identified in Incyte Clone 1824469 from the gallbladder tumor cDNA library (GBLADTUT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:113, was derived from Incyte Clones 1664262 (BRSTNOT09), 1733422 (BRSTTUT08), 1824469 (GBLADTUT01), 2057044 (BEPINOT01), and 2449822 (ENDANOT01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:36. SIGP-36 is 286 amino acids in length and has one potential N-glycosylation site at N271; four potential casein kinase II phosphorylation sites at S50, S192, T230, and T251; and five potential protein kinase C phosphorylation sites at T29, T41, S50, T160, and T273. SIGP-36 shares 24% identity with the Mycobacterium tuberculosis protein encoded by MTCI237.14c (GI 2052134). The fragment of SEQ ID NO:113 from about nucleotide 415 to about nucleotide 468 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, hematopoietic/immune, and neural cDNA libraries. Approximately 49% of these libraries are associated with cancer, 21% with immune response, and 21% with fetal/proliferating cells.

Nucleic acids encoding the SIGP-37 of the present invention were first identified in Incyte Clone 1864292 from the diseased prostate cDNA library (PROSNOT19) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:114, was derived from Incyte Clone 1864292 (PROSNOT19) and shotgun sequences SARA02195, SARA03070, SARA03675, and SATA02454.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:37. SIGP-37 is 404 amino acids in length and has one potential amidation site at V136; one potential cAMP- and cGMP-dependent protein kinase phosphorylation site at S66; twenty potential casein kinase II phosphorylation sites at S23, T27, T74, S110, S111, S118, T122, S143, S145, S205, S207, S218, S219, S220, T252, S254,

5

The state of the s

25

5

10

S328, S330, S385, and T393; and twelve potential protein kinase C phosphorylation sites at T27, S76, T81, S140, S161, S176, S229, T285, S309, S356, S367, and S398. SIGP-37 shares 18% identity with the <u>S. cerevisiae</u> protein encoded by SRP40, a weak suppressor of a mutant of the subunit AC40 of DNA-dependent RNA polymerases I and II (GI 295671).

The fragment of SEQ ID NO:114 f rom about nucleotide 193 to about nucleotide 222 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, cardiovascular, and hematopoietic/immune cDNA libraries. Approximately 75% of these libraries are associated with cancer and 25% with immune response.

Nucleic acids encoding the SIGP-38 of the present invention were first identified in Incyte Clone 1866437 from the human promonocyte cell line cDNA library (THP1NOT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:115, was derived from Incyte Clones 817970 (OVARTUT01), 825684 (PROSNOT06), 1866437 (THP1NOT01), 2190170 (PROSNOT26), and 3137972 (SMCCNOT02).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:38. SIGP-38 is 405 amino acids in length and has one potential N-glycosylation site at N378; one potential cAMP- and cGMP-phosphorylation site at S332; nine potential casein kinase II phosphorylation sites at T34, S51, T77, S107, S158, S264, T266, S296, and S332; and one potential protein kinase C phosphorylation site at S68. The fragment of SEQ ID NO:115 from about nucleotide 85 to about nucleotide 144 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, hematopoietic/immune, neural, and developmental cDNA libraries. Approximately 37% of these libraries are associated with cancer, 33% with immune response, and 22% with fetal/proliferating cells.

Nucleic acids encoding the SIGP-39 of the present invention were first identified in Incyte Clone 1871375 from the leg skin erythema nodosum cDNA library (SKINBIT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:116, was derived from Incyte Clones 1428052 (SINTBST01), 1871375 (SKINBIT01), and 3210563 (BLADNOT08).

5

10

## PF-0459 US

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:39. SIGP-39 is 177 amino acids in length and has one potential casein kinase II phosphorylation site at S133; one potential glycosaminoglycan attachment site at S28GGG; and four potential protein kinase C phosphorylation sites at S44, S82, S115, and T148. SIGP-39 contains a signature sequence shared by the binding domains of receptors for lymphokines, hematopoietic growth factors and growth hormone-related molecules at S52RWSLWS. The fragment of SEQ ID NO:116 encoding the sequence surrounding the receptor binding domain signature from about nucleotide 190 to about nucleotide 249 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, cardiovascular, gastrointestinal, and developmental cDNA libraries. Approximately 44% of these libraries are associated with cancer and 19% with immune response.

Nucleic acids encoding the SIGP-40 of the present invention were first identified in Incyte Clone 1880830 from the leukocyte cDNA library (LEUKNOT03) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:117, was derived from Incyte Clones 361577 (PROSNOT01); 2113591 (BRAITUT03); 1880830 (LEUKNOT03) and shotgun sequences SATA03292 and SATA00377.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:40. SIGP-40 is 197 amino acids in length and has a potential cAMP- and cGMP-dependent protein kinase phosphorylation site at S121; and four potential protein kinase C phosphorylation sites at T3, S57, T107, and T153. SIGP-40 shares 15% identity with the Arabidopsis thaliana zinc-finger protein Lsd1 (GI 1872521). The fragment of SEQ ID NO:117 from about nucleotide 567 to about nucleotide 621 is useful for hybridization. Northern analysis shows the expression of this sequence in neural and reproductive cDNA libraries. Approximately 49% of these libraries are associated with neoplastic disorders, 24% with immune response, and 16% with fetal development.

Nucleic acids encoding the SIGP-41 of the present invention were first identified in Incyte Clone 1905325 from the ovary cDNA library (OVARNOT07) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:118, was

and and that the set that the set that the set the set of the set

25

5

10

derived from Incyte Clones 1905325 (OVARNOT07); 621454 (PGANNOT01); 621326 (PGANNOT01); 1264490 (SYNORAT05); 487357 (HNT2AGT01); 773311 (COLNCRT01); and shotgun sequence SATA03582.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:41. SIGP-41 is 302 amino acids in length and has two potential N-glycosylation sites at N80 and N252; three potential casein kinase II phosphorylation sites at S46, T58, and S143; and four potential protein kinase C phosphorylation sites at T58, S62, T147, and S300. SIGP-41 shares 27% identity with human necdin-related protein (GI 1754971). The fragment of SEQ ID NO:118 from about nucleotide 1701 to about nucleotide 1800 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, neural, and gastrointestinal cDNA libraries. Approximately 51% of these libraries are associated with neoplastic disorders and 20% with immune response, and 18% with fetal development.

Nucleic acids encoding the SIGP-42 of the present invention were first identified in Incyte Clone 1919931 from the breast tumor cDNA library (BRSTTUT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:119, was derived from Incyte Clones 1919931 (BRSTTUT01) and shotgun sequences SATA02529, SATA01526 and SATA00892.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:42. SIGP-42 is 164 amino acids in length and has one potential casein kinase II phosphorylation site at T68; and two potential protein kinase C phosphorylation sites at T81 and S85. SIGP-42 shares 12% identity with human chemokine receptor (GI 2104517). The fragment of SEQ ID NO:119 from about nucleotide 585 to about nucleotide 630 is useful for hybridization. Northern analysis shows the expression of this sequence in hematopoietic/immune, reproductive, and neural cDNA libraries. Approximately 50% of these libraries are associated with neoplastic disorders and 38% with immune response.

Nucleic acids encoding the SIGP-43 of the present invention were first identified in Incyte Clone 1969426 from the breast tissue cDNA library (BRSTNOT04) using a

5

10

computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:120, was derived from Incyte Clones 1969426 (BRSTNOT04), 2373191 (ADRENOT07), 1225516 (COLNTUT02), 1555912 (BLADTUT04), 1449240 (PLACNOT02), and shotgun sequences SAZA01457 and SAZA00207.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:43. SIGP-43 is 235 amino acids in length and has one potential N-glycosylation site at N146; one potential glycosaminoglycan attachment site at S82; and four potential protein kinase C phosphorylation sites at T16, T43, S228, and S231. The fragment of SEQ ID NO:120 from about nucleotide 243 to about nucleotide 282 is useful for hybridization. Northern analysis shows the expression of this sequence in neural, reproductive, hematopoietic/immune, cardiovascular, gastrointestinal, and muscle cDNA libraries. Approximately 46% of these libraries are associated with neoplastic disorders and 28% with immune response.

Nucleic acids encoding the SIGP-44 of the present invention were first identified in Incyte Clone 1969948 from the umbilical cord cDNA library (UCMCL5T01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:121, was derived from Incyte Clones 1969948 (UCMCL5T01) and shotgun sequences SATA01513 and SATA00507.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:44. SIGP-44 is 203 amino acids in length and has three potential casein kinase II phosphorylation sites at T23, S114, and S120; one potential protein kinase C phosphorylation site at T105; and one potential tyrosine kinase phosphorylation site at Y47. The fragment of SEQ ID NO:121 from about nucleotide 162 to about nucleotide 216 is useful for hybridization. Northern analysis shows the expression of this sequence in gastrointestinal, hematopoietic/immune, reproductive, and cardiovascular cDNA libraries. Approximately 35% of these libraries are associated with neoplastic disorders and 24% with immune response.

Nucleic acids encoding the SIGP-45 of the present invention were first identified in Incyte Clone 1988911 from the lung cDNA library (LUNGAST01) using a computer

10

15 mm mm 1.4 mm 1.5 mm H. H. WA

20

25

search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:122, was derived from Incyte Clones 1988911 (LUNGAST01), 860576 (BRAITUT03), 3188894 (THYMNON04), 1466606 (PANCTUT02), 1920945 (BRSTTUT01), 1502970 (BRAITUT07), and shotgun sequence SAZC00040.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:45. SIGP-45 is 359 amino acids in length and has nine potential casein kinase II phosphorylation sites at S34, S47, S115, T120, T141, S157, S182, S214, and S331; three potential protein kinase C phosphorylation sites at S34, T259, and \$325; and one potential tyrosine kinase phosphorylation site at Y241. SIGP-45 shares 16% identity with rat myosin heavy chain (GI 56649). The fragment of SEQ ID NO:122 from about nucleotide 477 to about nucleotide 558 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, hematopoietic/immune, gastrointestinal, and cardiovascular cDNA libraries. Approximately 47% of these libraries are associated with neoplastic disorders, 33% with immune response, and 20% with fetal development.

Nucleic acids encoding the SIGP-46 of the present invention were first identified in Incyte Clone 2061561 from the ovary cDNA library (OVARNOT03) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:123, was derived from Incyte Clones 2061561 (OVARNOT03), 2208104 (SINTFET03), 2058750 (OVARNOT03), and shotgun sequences SAZA00915, SAZA00150, and SAZA00799.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:46. SIGP-46 is 150 amino acids in length and has two potential amidation sites at F57 and W74; one potential cAMP- and cGMP-dependent protein kinase phosphorylation site at T62; two potential casein kinase II phosphorylation sites at T101 and T110; and two potential protein kinase C phosphorylation sites at T28 and T97. The fragment of SEQ ID NO:123 from about nucleotide 82 to about nucleotide 168 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, neural, gastrointestinal, and cardiovascular cDNA libraries. Approximately 54% of these libraries are associated with neoplastic disorders and 22% with immune response.

5

10

## PF-0459 US

Nucleic acids encoding the SIGP-47 of the present invention were first identified in Incyte Clone 2084489 from the pancreas cDNA library (PANCNOT04) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:124, was derived from Incyte Clones 2084489 (PANCNOT04) and shotgun sequences SAJA00837, SAJA00793, SAJA01402, SAJA01533, and SAJA01490.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:47. SIGP-47 is 402 amino acids in length and has one potential N-glycosylation site at N191; seven potential cAMP- and cGMP-dependent protein kinase phosphorylation sites at S22, S23, T80, S81, S202, S248, and S382; twenty-two potential casein kinase II phosphorylation sites at S8, S35, S56, S107, T152, S166, S170, S202, S206, S208, T212, S214, S216, T244, S252, S256, T264, T287, S288, T327, S362, S387; ten potential protein kinase C phosphorylation sites at S16, S116, S140, T180, S193, S194, T236, T244, S252, and S387; and one potential tyrosine kinase phosphorylation site at Y361. SIGP-47 shares 28% identity with an A. thaliana protein of unknown function (GI 2262136). The most conserved region, residues 296 to 386 of SIGP-47, shares 70% identity with residues 299 to 386 of the A. thaliana protein. In addition, the potential amidation site at A314 in SIGP-47 is conserved as one potential amidation site at Q317 in the A. thaliana protein; and four potential protein kinase C or cAMP- and cGMP dependent protein kinase phosphorylation sites at S193, T236, S252 and Y361 in SIGP-47 are conserved as potential phosphorylation sites at S165, S219, T247, and Y364 respectively in the A. thaliana protein. The fragment of SEQ ID NO:124 from about nucleotide 468 to about nucleotide 531 is useful for hybridization. Northern analysis shows the expression of this sequence in neural, gastrointestinal and cardiovascular cDNA libraries. Approximately 50% of these libraries are associated with neoplastic disorders and 20% with trauma.

Nucleic acids encoding the SIGP-48 of the present invention were first identified in Incyte Clone 2203226 from the fetal spleen cDNA library (SPLNFET02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:125, was derived from Incyte Clones 2203226 (SPLNFET02), 2215960 (SINTFET03), 1291348

5

10

PF-0459 US

(BRAINOT11), 1874915 (LEUKNOT02), and 275828 (TESTNOT03).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:48. SIGP-48 is 311 amino acids in length and has one potential amidation site at V117; one potential casein kinase II phosphorylation site at T215; and three potential protein kinase C phosphorylation sites at T13, S18, and T263. SIGP-48 shares 32% identity with a human putative Rab5 interacting protein (GI 1911776). The fragment of SEQ ID NO:125 from about nucleotide 747 to about nucleotide 846 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, cardiovascular, neural, and gastrointestinal cDNA libraries. Approximately 44% of these libraries are associated with neoplastic disorders, 30% with fetal/proliferative cells and tissues, and 23% with immune response.

Nucleic acids encoding the SIGP-49 of the present invention were first identified in Incyte Clone 2232884 from the prostate cDNA library (PROSNOT16) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:126, was derived from Incyte Clones 2232884 (PROSNOT16), 2728528 (OVARTUT05), 2232884 (PROSNOT16), and shotgun sequences SASA00238 and SASA00455.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:49. SIGP-49 is 316 amino acids in length and has one potential N-glycosylation site at N140; five potential casein kinase II phosphorylation sites at S3, T8, S29, S85, and T198; and two potential protein kinase C phosphorylation sites at T28 and S60. The fragment of SEQ ID NO:126 from about nucleotide 180 to about nucleotide 279 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, urologic, and neural cDNA libraries. Approximately 77% of these libraries are associated with neoplastic disorders.

Nucleic acids encoding the SIGP-50 of the present invention were first identified in Incyte Clone 2328134 from the colon cDNA library (COLNNOT11) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:127, was derived from Incyte Clones 2328134 (COLNNOT11), 1870180 (SKINBIT01), 081403 (SYNORAB01), and 851547 (NGANNOT01).

30

5

.

10

### PF-0459 US

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:50. SIGP-50 is 346 amino acids in length and has two potential cAMP- and cGMP-dependent protein kinase phosphorylation sites at residues S43 and S217; one potential casein kinase II phosphorylation site at residue T96; and five potential protein kinase C phosphorylation sites at residues T2, T15, T39, T247, and S301. SIGP-50 shares 33% identity with the human putative rab5-interacting protein (GI 1911776) and the casein kinase II phosphorylation site at residue T96. The fragment of SEQ ID NO:127 encoding the potential extracellular ligand binding domain from about nucleotide 16 to about nucleotide 76 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, cardiovascular, and neural cDNA libraries. Approximately 44% of these libraries are associated with cancer, 28% are associated with immune response, and 20% with fetal disorders.

Nucleic acids encoding the SIGP-51 of the present invention were first identified in Incyte Clone 2382718 from the pancreatic cDNA library (ISLTNOT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:128, was derived from Incyte Clones 2382718 (ISLTNOT01), 3472492 (LUNGNOT27), 014756 (THP1PLB01), 1731885 (BRSTTUT08), 1889866 (BLADTUT07), and 1447744 (PLACNOT02).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:51. SIGP-51 is 299 amino acids in length and has one potential N-glycosylation site at residue N185; one cAMP- and cGMP-dependent protein kinase phosphorylation site at T273; nine potential casein kinase II phosphorylation sites at S34, S82, T100, S118, T152, S154, T193, S203, and S287; eight potential protein kinase C phosphorylation sites at S57, T69, T95, S179, T269, S274, S275, and S284; and a potential signal peptide sequence from M1 to G27. SIGP-51 shares 26% identity with a human antigen precursor protein (GI 1814277); the protein kinase C phosphorylation sites at residues S57 and T69; and the casein kinase II phosphorylation site at residue T100. The fragment of SEQ ID NO:128 encoding the potential extracellular ligand binding domain from about nucleotide 88 to about nucleotide 148 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal,

30

5

10

and cardiovascular cDNA libraries. Approximately 48% of these libraries are associated with cancer, 29% are associated with immune response, and 20% with fetal disorders.

Nucleic acids encoding the SIGP-52 of the present invention were first identified in Incyte Clone 2452208 from the cardiovascular cDNA library (ENDANOT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:129, was derived from Incyte Clones 2452280 (ENDANOT01), 1505094 (BRAITUT07), 1521239 (BLADTUT04), and 1309844 (COLNFET02).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:52. SIGP-52 is 351 amino acids in length and has two potential N-glycosylation sites at N241 and N337; two potential cAMP- and cGMP-dependent protein kinase phosphorylation sites at S201 and T318; six potential casein kinase II phosphorylation sites at S9, S136, T162, T252, S270, and S302; eight potential protein kinase C phosphorylation sites at T25, S34, T37, S64, S87, S112, S141, and S322; and one potential cell attachment sequence at R280GD. The fragment of SEQ ID NO:129 encoding the potential extracellular ligand binding domain from about nucleotide 97 to about nucleotide 157 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, cardiovascular, and neural cDNA libraries. Approximately 33% of these libraries are associated with cancer, 33% are associated with immune response, and 26% with fetal disorders.

Nucleic acids encoding the SIGP-53 of the present invention were first identified in Incyte Clone 2457825 from the aortic endothelial cell cDNA library (ENDANOT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:130, was derived from Incyte Clone 2457825 (ENDANOT01) and shotgun sequences SASA00641, SASA02817, SASA01973, SASA03121, SASA01350, and SASA00693.

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:53. SIGP-53 is 662 amino acids in length and has three potential cAMP- and cGMP-dependent protein kinase phosphorylation sites at S555, S578, and S652; ten potential casein kinase II phosphorylation sites at S67, T151, T215, S241, S470, S471, S482, S556, T589, and T618; one potential leucine zipper pattern from L572 to L593; four potential protein kinase C phosphorylation sites at T2, T21, S80, and T503;

30

5

10

and one potential LIM domain signature site from C402 to L436. SIGP-53 shares 10% identity with the <u>C. elegans</u> protein encoded by W04D2.1 (GI 1418625); and the casein kinase II phosphorylation site at residue S241. The fragment of SEQ ID NO:130 encoding the potential extracellular ligand binding domain from about nucleotide 88 to about nucleotide 148 is useful for hybridization. Northern analysis shows the expression of this sequence in hematopoietic, gastrointestinal, reproductive, and cardiovascular cDNA libraries. Approximately 43% of these libraries are associated with cancer, 35% are associated with immune response, and 22% with fetal disorders.

Nucleic acids encoding the SIGP-54 of the present invention were first identified in Incyte Clone 2470740 from the hematopoietic cDNA library (THP1NOT03) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:131, was derived from Incyte Clone 2470740 (THP1NOT03).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:54. SIGP-54 is 115 amino acids in length and has one potential protein kinase C phosphorylation site at S85; and one potential insulin family signature site from C23 to C37. The fragment of SEQ ID NO:131 encoding the potential extracellular ligand binding domain from about nucleotide 151 to about nucleotide 211 is useful for hybridization. Northern analysis shows the expression of this sequence in neural and developmental cDNA libraries. Approximately 33% of these libraries are associated with cancer and 33% are associated with fetal disorders.

Nucleic acids encoding the SIGP-55 of the present invention were first identified in Incyte Clone 2479092 from the aortic endothelial cell cDNA library (SMCANOT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:132, was derived from Incyte Clone 2479092 (SMCANOT01) and 1981954 (LUNGTUT03).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:55. SIGP-55 is 157 amino acids in length and has one potential casein kinase II phosphorylation site at S31; one potential tyrosine kinase phosphorylation site at K150; and a potential signal peptide sequence from M1 to A26. The fragment of SEQ ID NO:132 encoding the potential extracellular ligand binding

30

5

10

## PF-0459 US

domain from about nucleotide 97 to about nucleotide 157 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, hematopoietic, and urologic cDNA libraries. Approximately 47% of these libraries are associated with cancer and 29% with immune response.

Nucleic acids encoding the SIGP-56 of the present invention were first identified in Incyte Clone 2480544 from the aortic smooth muscle cell cDNA library (SMCANOT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:133, was derived from Incyte Clones 2480544 (SMCANOT01), 2472409 (THP1NOT03), 1516031 (PANCTUT01), 855817 (NGANNOT01), 1865287 (PROSNOT19), and 677835 (CRBLNOT01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:56. SIGP-56 is 197 amino acids in length and has one potential N glycosylation site at N38; one potential casein kinase II phosphorylation site at S123; two potential protein kinase C phosphorylation sites at T71 and S82; and a potential signal peptide sequence from M1 to A27. SIGP-56 shares 15% identity with a Phaseolus vulgaris protein involved in the stress response (GI 169345) and shows conservation of proline and tyrosine residues in the C-terminal region. The fragment of SEQ ID NO:133 from about nucleotide 125 to about nucleotide 160 is useful for hybridization. Northern analysis shows the expression of this sequence in neural, reproductive, and cardiovascular cDNA libraries. Approximately 49% of these libraries are associated with neoplastic disorders and 14% with immune response.

Nucleic acids encoding the SIGP-57 of the present invention were first identified in Incyte Clone 2518547 from the brain tumor cDNA library (BRAITUT21) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:134, was derived from Incyte Clones 2518547 (BRAITUT21), 1509622 (LUNGNOT14), 1562945 (SPLNNOT04), 1640136 (UTRSNOT06), and 1432014 (BEPINON01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:57. SIGP-57 is 245 amino acids in length and has one potential casein kinase II phosphorylation site at S27; and two potential protein kinase C phosphorylation sites at S5 and T229. SIGP-57 shares 36% identity with a human protein

30

5

10

## PF-0459 US

that binds a regulatory element of the c-myc gene (GI 33969). In addition, the potential protein kinase C phosphorylation site at T229 is conserved as a potential protein kinase A phosphorylation site at S176 in the human protein. The fragment of SEQ ID NO:134 from about nucleotide 742 to about nucleotide 775 is useful for hybridization. Northern analysis shows the expression of this sequence in hematopoietic, reproductive, and neural cDNA libraries. Approximately 50% of these libraries are associated with neoplastic disorders and 28% with immune response.

Nucleic acids encoding the SIGP-58 of the present invention were first identified in Incyte Clone 2530650 from the gallbladder cDNA library (GBLANOT02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:135, was derived from Incyte Clones 2530650 (GBLANOT02), 2617724 (GBLANOT01), 3105644 (BRSTTUT15), 2903466 (DRGCNOT01), 1545010 (PROSTUT04), 2313837 (NGANNOT01), 1804413 (SINTNOT13), 3207379 (PENCNOT03), 2347051 (TESTTUT02), 2602493 (UTRSNOT10), 1259341 (MENITUT03), and 81943 (SYNORAB01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:58. SIGP-58 is 310 amino acids in length and has one potential N glycosylation site at N206; one potential cAMP- and cGMP-dependent protein kinase phosphorylation site at T97; five potential casein kinase II phosphorylation sites at S62, S156, S214, S222, and T274; five potential protein kinase C phosphorylation sites at T150, T167, T208, T265, and S273; one potential tyrosine kinase phosphorylation site at Y96; one thyroglobulin type-1 repeat signature from F109 to G143; and a potential signal peptide sequence from M1 to A21. SIGP-58 shares 18% identity with bovine thyroglobulin (GI 2204111) and 46% identity between F109 and G143, the thyroglobulin type-1 repeat signature. The fragment of SEQ ID NO:135 from about nucleotide 92 to about nucleotide 127 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive and cardiovascular cDNA libraries. Approximately 67% of these libraries are associated with neoplastic disorders and 19% with immune response.

Nucleic acids encoding the SIGP-59 of the present invention were first identified in Incyte Clone 2652271 from the thymus cDNA library (THYMNOT04) using a computer

30

5

10

### PF-0459 US

search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:136, was derived from Incyte Clones 2652271 (THYMNOT04), 2742813 (BRSTTUT14), 763431 (BRAITUT02), 1272403 (TESTTUT02), 1240531 (LUNGNOT03), and 1318448 (BLADNOT04).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:59. SIGP-59 is 256 amino acids in length and has three potential N glycosylation sites at N76, N106, and N212; three potential casein kinase II phosphorylation sites at T46, S188, and T204; two potential protein kinase C phosphorylation sites at S130 and S221; two potential ribonuclease T2 family histidine active sites from W62 to P69 and from F110 to C121; and a potential signal peptide sequence from M1 to A24. SIGP-59 shares 24% identity with Solanum lycopersicum ribonuclease LE (GI 895855); 80% identity between W62 and P75, one of the two ribonuclease T2 family histidine active sites; and 92% identity between F110 and C121, the second of the two ribonuclease T2 family histidine active sites. The fragment of SEQ ID NO:136 from about nucleotide 462 to about nucleotide 494 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, hematopoietic, and gastrointestinal cDNA libraries. Approximately 53% of these libraries are associated with neoplastic disorders and 28% with immune response.

Nucleic acids encoding the SIGP-60 of the present invention were first identified in Incyte Clone 2746976 from the lung tumor cDNA library (LUNGTUT11) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:137, was derived from Incyte Clones 2746976 (LUNGTUT11), 488049 (HNT2AGT01), 1907738 (CONNTUT01), 782645 (MYOMNOT01), and 823864 (PROSNOT06).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:60. SIGP-60 is 160 amino acids in length and has one potential cAMP- and cGMP-dependent protein kinase phosphorylation site at T31; four potential casein kinase II phosphorylation sites at S23, S47, S96, and S152; four potential protein kinase C phosphorylation sites at S23, T125, S126, and T149; and a clathrin adaptor complex small chain signature from I56 to F66. SIGP-60 shares 84% identity with mouse clathrin-associated protein 19 (GI 191983) and 91% identity with the clathrin adaptor complex small

30

5

10

# PF-0459 US

chain signature between I56 and F66. In addition, all potential casein kinase II and protein kinase C phosphorylation sites are conserved between SIGP-60 and the mouse protein. The fragments of SEQ ID NO:137 from about nucleotide 144 to about nucleotide 170 and from about nucleotide 495 to about nucleotide 521 are useful for hybridization. Northern analysis shows the expression of this sequence in hematopoietic, cardiovascular, and reproductive cDNA libraries. Approximately 39% of these libraries are associated with neoplastic disorders and 39% with immune response.

Nucleic acids encoding the SIGP-61 of the present invention were first identified in Incyte Clone 2753496 from the THP-1 promonocyte cDNA library (THP1AZS08) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:138, was derived from Incyte Clones 2753496 (THP1AZS08), 2642512 (LUNGTUT08), 1367244 (SCORNON02), 474458 (MMLR1DT01), 1349777 (LATRTUT02), 1380831 (BRAITUT08), and 832934 (PROSTUT04).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:61. SIGP-61 is 341 amino acids in length and has one potential N glycosylation site at N66; four potential casein kinase II phosphorylation sites at T157, T207, S296, and S335; two potential protein kinase C phosphorylation sites at S159 and S296; and one potential tyrosine kinase phosphorylation site at Y184. SIGP-61 shares 17% identity with Schizosaccharomyces pombe BEM46, a protein involved in cell polarity (GI 987286) and the potential phosphorylation sites at T157 and S296. The fragment of SEQ ID NO:138 from about nucleotide 79 to about nucleotide 114 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, and neural cDNA libraries. Approximately 52% of these libraries are associated with neoplastic disorders and 25% with immune response.

Nucleic acids encoding the SIGP-62 of the present invention were first identified in Incyte Clone 2781553 from the ovarian tumor cDNA library (OVARTUT03) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:139, was derived from Incyte Clones 2781553 (OVARTUT03), 1413079 (BRAINOT12), 894971 (BRSTNOT05), 2696043 (UTRSNOT12), 1267806 (BRAINOT09), 1961608 (BRSTNOT04), 1755817 (LIVRTUT01), 1793882 (PROSTUT05), 1251515

10

PF-0459 US

(LUNGFET03), 1560984 (SPLNNOT04), and 1872574 (LEUKNOT02).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:62. SIGP-62 is 430 amino acids in length and has one potential cAMP- and cGMP-dependent protein kinase phosphorylation site at S387; thirteen potential casein kinase II phosphorylation sites at S182, S214, S235, T248, S258, T266, T275, T294, S313, T356, S387, T404, and S413; six potential protein kinase C phosphorylation sites at T71, S168, S235, S306, T356, and S374; and a mitochondrial energy transfer protein signature from P114 to L122. Northern analysis shows the expression of this sequence in reproductive, neural, and hematopoietic cDNA libraries. Approximately 47% of these libraries are associated with neoplastic disorders and 19% with immune response.

Nucleic acids encoding the SIGP-63 of the present invention were first identified in Incyte Clone 2821925 from the adrenal tumor cDNA library (ADRETUT06) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:140, was derived from Incyte Clones 2821925 (ADRETUT06), 933799 (CERVNOT01), and 136467 (SYNORAB01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:63. SIGP-63 is 143 amino acids in length and has one potential cAMP- and cGMP-dependent protein kinase phosphorylation site at S109; three potential casein kinase II phosphorylation sites at S36, S80, and T84; five potential protein kinase C phosphorylation sites at T31, T55, T70, S109, and T122; and a potential signal peptide sequence from M1 to A21. Northern analysis shows the expression of this sequence in reproductive, musculoskeletal and cardiovascular cDNA libraries. Approximately 50% of these libraries are associated with neoplastic disorders and 27% with immune response.

Nucleic acids encoding the SIGP-64 of the present invention were first identified in Incyte Clone 2879068 from the uterine tumor cDNA library (UTRSTUT05) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:141, was derived from Incyte Clones 2879068 (UTRSTUT05), 2910155 (KIDNTUT15), 488673 (HNT2AGT01), 1285407 (COLNNOT16), 1415890 (BRAINOT12), 1352662 (LATRTUT02), 41046 (TBLYNOT01), and 2686554 (LUNGNOT23).

In one embodiment, the invention encompasses a polypeptide comprising the amino

30

25

30

10

# PF-0459 US

acid sequence of SEQ ID NO:64. SIGP-64 is 301 amino acids in length and has two potential N glycosylation sites at N20 and N251; five potential casein kinase II phosphorylation sites at S8, S41, T125, T161, and T163; five potential protein kinase C phosphorylation sites at T40, S41, T59, T66, and S181; one potential tyrosine kinase phosphorylation site at Y176; one potential glycosaminoglycan attachment site at S253; and two putative RNP-1 RNA-binding signatures from R70 to F77 and from R155 to Y162. SIGP-64 shares 59% identity with human heterogeneous nuclear ribonucleoprotein D (GI 870749); 100% identity between R70 and F77, one of the two RNP-1 RNA-binding signatures; and 89% identity between R155 and Y162, the second of the two RNP-1 RNA-binding signatures. In addition, eight potential phosphorylation sites are conserved between SIGP-64 and the human ribonucleoprotein. The fragments of SEQ ID NO:141 from about nucleotide 207 to about nucleotide 248 and from about nucleotide 726 to about nucleotide 752 are useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, neural, hematopoietic, and gastrointestinal cDNA libraries. Approximately 48% of these libraries are associated with neoplastic disorders and 24% with immune response.

Nucleic acids encoding the SIGP-65 of the present invention were first identified in Incyte Clone 2886757 from the small intestine cDNA library (SINJNOT02) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:142, was derived from Incyte Clones 2886757 (SINJNOT02), 2230747 (PROSNOT16), and 899432 (BRSTTUT03).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:65. SIGP-65 is 233 amino acids in length and has two potential N-glycosylation sites at N82 and N196; one potential casein kinase II phosphorylation site at S170; and two potential protein kinase C phosphorylation sites at S102 and T134. SIGP-65 shares 22% identity with S. cerevisiae protein encoded by YOL135c (GI 1420026), and the potential casein kinase II phosphorylation site at S170 is conserved between the two proteins. The fragment of SEQ ID NO:142 from about nucleotide 99 to about nucleotide 137 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, cardiovascular, and gastrointestinal cDNA libraries. Approximately 59% of these libraries are associated with neoplastic disorders.

30

5

10

## PF-0459 US

Nucleic acids encoding the SIGP-66 of the present invention were first identified in Incyte Clone 2964329 from the cervical spinal cord cDNA library (SCORNOT04) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:143, was derived from Incyte Clones 2964329, (SCORNOT04), 1274814 (TESTTUT02), 746049 (BRAITUT01), 1395667 (THYRNOT03), 1362944 (LUNGNOT12), and 2589 (HMC1NOT01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:66. SIGP-66 is 354 amino acids in length and has one potential cAMP- and cGMP-dependent protein kinase phosphorylation site at S346; two potential casein kinase II phosphorylation sites at S164 and T180; six potential protein kinase C phosphorylation sites at S43, S135, S150, S164, S172, and S201; and one potential tyrosine kinase phosphorylation site at Y182. SIGP-66 shares 12% identity with S. cerevisiae mitochondrial internal membrane carrier protein (GI 311667). In addition, one potential protein kinase C site is conserved between these molecules. The fragment of SEQ ID NO:143 from about nucleotide 416 to about nucleotide 442 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, neural, hematopoietic/immune, gastrointestinal, and cardiovascular cDNA libraries. Approximately 46% of these libraries are associated with neoplastic disorders and 26% with immune response.

Nucleic acids encoding the SIGP-67 of the present invention were first identified in Incyte Clone 2965248 from the cervical spinal cord cDNA library (SCORNOT04) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:144, was derived from Incyte Clones 2965248 (SCORNOT04), 485746 (HNT2RAT01), 865684 (BRAITUT03), 1459157 (COLNFET02), 1597772 (BRAINOT14), 531430 (BRAINOT03), 725362 (SYNOOAT01), 1620429 (BRAITUT13), and 190305 (SYNORAB01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:67 SIGP-67 is 235 amino acids in length and has seven potential cAMP- and cGMP-dependent protein kinase phosphorylation sites at S50, T80, T98, T126, S135, S136, and T194; three potential casein kinase II phosphorylation sites at

30

5

10

S60, T80, and S81; six potential protein kinase C phosphorylation sites at S114, T119, T137, S142, S146, and S174; and a strathmin 1 family signature from P75 to E84. SIGP-67 shares 44% identity with human strathmin homolog SCG10/neuron-specific growth-associated protein in Alzheimer's disease (GI 1478503), and 71% identity between M1 and A107. In addition, one potential cAMP- and cGMP-dependent protein kinase phosphorylation site, one potential casein kinase II phosphorylation site, the strathmin 1 family signature, and the hydrophobic transmembrane domains are conserved between these molecules. TM1 extends from about L15 to about F25; and TM2, from about G196 to about P212. The fragments of SEQ ID NO:144 from about nucleotide 158 to about nucleotide 196 and from about nucleotide 614 to about nucleotide 643 are useful for hybridization. Northern analysis shows the expression of this sequence in neural, reproductive, gastrointestinal, and hematopoietic/immune cDNA libraries. Approximately 50% of these libraries are associated with neoplastic disorders and 19% with immune response.

Nucleic acids encoding the SIGP-68 of the present invention were first identified in Incyte Clone 3000534 from the Th2 T lymphocyte cDNA library (TLYMNOT06) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:145, was derived from Incyte Clones 3000534 (TLYMNOT06), 1830964 (THP1AZT01), 1329136 (PANCNOT07), and 2910083 (KIDNTUT15).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:68. SIGP-68 is 221 amino acids in length and has two potential casein kinase II phosphorylation sites at T31 and T70; one potential glycosaminoglycan attachment site at S62; three potential protein kinase C phosphorylation sites at T111, T146, and T199; and an endoplasmic reticulum targeting sequence at H218DEL. SIGP-68 shares 61% identity with the human stroma cell-derived secretory factor-2 (GI 1741868). In addition, one potential protein kinase C phosphorylation site and the hydrophobic transmembrane domains are conserved between these molecules. TM1 extends from about A10 to about G27; and TM2, from about T31 to about L45. The cysteines at C38, C92, C100, and C149 are conserved between both molecules. The fragments of SEQ ID NO:145 from about nucleotide 89 to about nucleotide 118 and from

30

5

10

about nucleotide 608 to about nucleotide 643 are useful for hybridization. Northern analysis shows the expression of this sequence in hematopoietic/immune, reproductive, cardiovascular, and gastrointestinal cDNA libraries. Approximately 41% of these libraries are associated with neoplastic disorders and 31% with immune response.

Nucleic acids encoding the SIGP-69 of the present invention were first identified in Incyte Clone 3046870 from the coronary artery cDNA library (HEAANOT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:146, was derived from Incyte Clones 3046870 (HEAANOT01), 2719210 (THYRNOT09), 581291 (SATPFI006), 1961256 (BRSTNOT04), 2226972 (SEMVNOT01), 2023351 (CONNNOT01), 1379008 (LUNGNOT10), and 1943136 (HIPONOT01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:69. SIGP-69 is 483 amino acids in length and has one potential N-glycosylation site at N178; ten potential casein kinase II phosphorylation sites at S16, S49, T60, T67, T92, T121, T170, T187, T250, and S431; and nine potential protein kinase C phosphorylation sites at S113, T170, T187, T194, S210, T265, S284, T355, and S431. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, cardiovascular, and neural cDNA libraries. Approximately 49% of these libraries are associated with neoplastic disorders and 24% with immune response.

Nucleic acids encoding the SIGP-70 of the present invention were first identified in Incyte Clone 3057669 from the pons cDNA library (PONSAZT01) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:147, was derived from Incyte Clones 3057669 (PONSAZT01), 548211 (BEPINOT01), 3702516 (PENCNOT07), 3581270 (293TF3T01), 495191 (HNT2NOT01), 2784427 (BRSTNOT13), 1515961 (PANCTUT01), 3552333 (SYNONOT01), 2838668 (DRGLNOT01), 14600680 (COLNFET02), and 285677 (EOSIHET02).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:70. SIGP-70 is 371 amino acids in length and has three potential N-glycosylation sites at N70, N125, and N362; eleven potential casein kinase II phosphorylation sites at T22, S66, S72, S73, S102, T160, T201, T215, T278, T285, and

30

5

10

S316; seven potential protein kinase C phosphorylation sites at S72, T79, S99, T127, S134, S257, and T299; and one protein kinase signature and profile from L188 to F200. Northern analysis shows the expression of this sequence in gastrointestinal, reproductive, and neural cDNA libraries. Approximately 54% of these libraries are associated with neoplastic disorders and 14% with immune response.

Nucleic acids encoding the SIGP-71 of the present invention were first identified in Incyte Clone 3088178 from the aorta cDNA library (HEAONOT03) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:148, was derived from Incyte Clones 3088178 (HEAONOT03), 589421 (UTRSNOT01), 2059958 (OVARNOT03), 1550631 (PROSNOT06), and 1271480 (TESTTUT02).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:71. SIGP-71 is 402 amino acids in length and has two potential N glycosylation sites at N13 and N366; two potential cAMP- and cGMP-dependent protein kinase phosphorylation sites at T50 and S51; five potential casein kinase II phosphorylation sites at T50, S51, S52, S56, and S246; one potential glycosaminoglycan attachment site at S247; eight potential protein kinase C phosphorylation sites at T45, T46, S224, S240, S259, T279, S338, and S376; one potential tyrosine kinase phosphorylation site at Y273; and one beta-transducin family Trp-Asp repeat signature from V243 to V257. SIGP-71 shares 22% identity with S. cerevisiae protein encoded by HRE594 (GI 498997; truncated sequence). In addition, one potential N-glycosylation site, and two potential casein kinase II phosphorylation sites are conserved between these molecules. The fragment of SEQ ID NO:148 from about nucleotide 725 to about nucleotide 766 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, neural, cardiovascular, and hematopoietic/immune cDNA libraries. Approximately 51% of these libraries are associated with neoplastic disorders and 23% with immune response.

Nucleic acids encoding the SIGP-72 of the present invention were first identified in Incyte Clone 3094321 from the breast cDNA library (BRSTNOT19) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:149, was derived from Incyte Clones 3094321 (BRSTNOT19), 2517422H1 (BRAITUT21), 2101110 (BRAITUT02), 1303603 (PLACNOT02), 2675275 (KIDNNOT19), 1988065

Line half be then then the man he may take the term that the term that the

20

25

30

5

10

(LUNGAST01), 34101 (THP1NOB01), 1815156 (PROSNOT20), 602724 (BRSTTUT01), and 1485067 (CORPNOT02).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:72. SIGP-72 is 640 amino acids in length and has four potential N-glycosylation sites at N295, N513, N568, and N619; two potential cAMP- and cGMP-dependent protein kinase phosphorylation sites at S239 and S507; sixteen potential casein kinase II phosphorylation sites at S42, T178, T220, S229, S239, T247, S289, S350, S372, S446, T463, S492, T580, S592, S604, and S625; nine potential protein kinase C phosphorylation sites at T150, T166, T174, S239, T328, S407, T451, S609, and S621; one potential tyrosine kinase phosphorylation site at Y265; and one cytochrome c family hemebinding site signature at C158YECHP. SIGP-72 shares 33% identity with an essential yeast ubiquitin-activating enzyme homolog (GI 793879). In addition, one potential Nglycosylation site, one potential casein kinase II phosphorylation site, and six potential protein kinase C phosphorylation sites are conserved between these molecules. The fragments of SEQ ID NO:149 from about nucleotide 382 to about nucleotide 423 and from about nucleotide 1087 to about nucleotide 1113 are useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, hematopoietic/immune, cardiovascular, and gastrointestinal cDNA libraries. Approximately 48% of these libraries are associated with neoplastic disorders and 24% with immune response.

Nucleic acids encoding the SIGP-73 of the present invention were first identified in Incyte Clone 3115936 from the lung cDNA library (LUNGTUT13) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:150, was derived from Incyte Clones 3115936 (LUNGTUT13) 2359411 (LUNGFET05), 2189762 (PROSNOT26), 1449756 (PLACNOT02), 541212 (LNODNOT02), 079364 (SYNORAB01), 864877 (BRAITUT03), 2697958 (UTRSNOT12), 1818830 (PROSNOT20), 1966765 (BRSTNOT04), 998279 (KIDNTUT01), 1961616 (BRSTNOT04), and 1431515 (BEPINON01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:73. SIGP-73 is 237 amino acids in length and has five potential casein kinase II phosphorylation sites at S43, S47, S72, S131, and T177; and

25

30

5

10

three potential protein kinase C phosphorylation sites at S39, S125, and T202. SIGP-73 shares 44% identity with t yeast Rer1p protein, which ensures correct localization of Sec12p integral membrane protein of the endoplasmic reticulum (GI 517174). In addition, the hydrophobic transmembrane domains are conserved among these molecules. TM1 extends from about A82 to about P126; and TM2, from about A166 to about M203. The fragment of SEQ ID NO:150 from about nucleotide 585 to about nucleotide 623 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, neural, cardiovascular, gastrointestinal, and hematopoietic/ immune cDNA libraries. Approximately 48% of these libraries are associated with neoplastic disorders and 24% with immune response.

Nucleic acids encoding the SIGP-74 of the present invention were first identified in Incyte Clone 3116522 from the lung cDNA library (LUNGTUT13) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:151, was derived from Incyte Clones 3116522 (LUNGTUT13), 2523149 (BRAITUT21), 1513583 (PANCTUT01), 834017 (PROSNOT07), 1631796 (COLNNOT19), 1502736 (BRAITUT07), and 78850 (SYNORAB01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:74. SIGP-74 is 432 amino acids in length and has three potential casein kinase II phosphorylation sites at S144, S257, and S317; three potential protein kinase C phosphorylation sites at T68, S231, and T372; and one potential tyrosine kinase phosphorylation site at Y240. SIGP-74 shares 28% identity with the human UDP-galactose transporter isoform (GI 1669560). In addition, one potential protein kinase C phosphorylation site and the hydrophobic transmembrane domains are conserved between these molecules. TM4 extends from about Q108 to about G127; TM5, from about S152 to about L173; TM6, from about K205 to about K228; TM7, from about T242 to about S257; TM8, from about T268 to about S283; TM9, from about A294 to about T328; and TM10, from about A338 to about V409. The fragment of SEQ ID NO:151 from about nucleotide 710 to about nucleotide 736 is useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, gastrointestinal, cardiovascular, hematopoietic/immune, and urologic cDNA libraries. Approximately 54% of these

30

5

10

## PF-0459 US

libraries are associated with neoplastic disorders and 25% with immune response.

Nucleic acids encoding the SIGP-75 of the present invention were first identified in Incyte Clone 3117184 from the lung cDNA library (LUNGTUT13) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:152, was derived from Incyte Clones 3117184 (LUNGTUT13), 2494724 (ADRETUT05), and 1922002 (BRSTTUT01).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:75. SIGP-75 is 252 amino acids in length and has one potential N-glycosylation site at N93; one potential cAMP- and cGMP-dependent protein kinase phosphorylation site at S179; one potential casein kinase II phosphorylation site at T189; and five potential protein kinase C phosphorylation sites at S95, S115, S123, T140, and T200. SIGP-75 shares 39% identity with C. elegans protein encoded by WO4D2.6 (GI 1418628). In addition, one potential N-glycosylation site, and three potential protein kinase C phosphorylation sites are conserved between the molecules. The fragment of SEQ ID NO:152 from about nucleotide 567 to about nucleotide 593 is useful for hybridization. Northern analysis shows the expression of this sequence in cardiovascular, gastrointestinal, hematopoietic/immune, and reproductive cDNA libraries. Approximately 50% of these libraries are associated with neoplastic disorders and 20% with immune response.

Nucleic acids encoding the SIGP-76 of the present invention were first identified in Incyte Clone 3125156 from the lymph node cDNA library (LNODNOT05) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:153, was derived from Incyte Clones 3125156 (LNODNOT05), 1417459 (BRAINOT12), 1567861 (UTRSNOT05), 154233 (THP1PLB02), 872652 (LUNGAST01), 2525803 (BRAITUT21), and 1209172 (BRSTNOT02).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:76. SIGP-76 is 523 amino acids in length and has one potential N glycosylation sites at N186; nine potential casein kinase II phosphorylation sites at S63, T85, S179, S188, T210, S231, T269, T295, and S474; one potential glycosaminoglycan attachment site at S335; ten potential protein kinase C phosphorylation sites at T9, S159, S172, S179, T246, S263, S283, S416, S447, and S498; two potential tyrosine kinase

25

30

5

10

phosphorylation sites at Y106 and Y170; and one tyrosine specific protein phosphatase active site at V331. SIGP-76 shares 21% identity with human T-cell protein tyrosine phosphatase (GI 804750), the N186 glycosylation site, the phosphorylation sites at S179, S188, T210, T246, S263, T295, S416, and Y170; and 50% identity between P324 and F344, the region of the tyrosine specific protein phosphatase active site. The fragments of SEQ ID NO:153 from about nucleotide 64 to about nucleotide 183 and from about nucleotide 1087 to about nucleotide 1119 are useful for hybridization. Northern analysis shows the expression of this sequence in neural, reproductive, and gastrointestinal cDNA libraries. Approximately 55% of these libraries are associated with neoplastic disorders and 22% with immune response.

Nucleic acids encoding the SIGP-77 of the present invention were first identified in Incyte Clone 3129120 from the lung tumor cDNA library (LUNGTUT12) using a computer search for amino acid sequence alignments. A consensus sequence, SEQ ID NO:154, was derived from Incyte Clones 3129120 (LUNGTUT12), 3744590 (THYMNOT08), 1512939 (PANCTUT01), 3220539 (COLNNON03), 1435889 (PANCNOT08), 1452745 (PENITUT01), 874548 (LUNGAST01), 1524326 (UCMCL5T01), and 811239 (LUNGNOT04).

In one embodiment, the invention encompasses a polypeptide comprising the amino acid sequence of SEQ ID NO:77. SIGP-77 is 621 amino acids in length and has two potential N glycosylation sites at N203 and N517; one potential protein kinase A or G phosphorylation site at S84; five potential casein kinase II phosphorylation sites at T45, T185, T233, T278, and S573; seven potential protein kinase C phosphorylation sites at T45, T95, S109, S299, T318, S324, and T482; and one potential leucine zipper motif from L332 to L353. SIGP-77 shares 27% identity and the phosphorylation site at T318 with S. cerevisiae membrane protein important for endocytosis (GI 1256890). The fragments of SEQ ID NO:154 from about nucleotide 64 to about nucleotide 183 and from about nucleotide 1087 to about nucleotide 1119 are useful for hybridization. Northern analysis shows the expression of this sequence in reproductive, neural, gastrointestinal, and cardiovascular cDNA libraries. Approximately 53% of these libraries are associated with neoplastic disorders and 17% with immune response.

The invention also encompasses SIGP variants. A preferred SIGP variant is one which

20

25

30

5

10

### PF-0459 US

has at least about 80%, more preferably at least about 90%, and most preferably at least about 95% amino acid sequence identity to the SIGP amino acid sequence, and which contains at least one functional or structural characteristic of SIGP.

The invention also encompasses polynucleotides which encode SIGP. Accordingly, any nucleic acid sequence which encodes the amino acid sequence of SIGP can be used to produce recombinant molecules which express SIGP. In a particular embodiment, the invention encompasses a polynucleotide consisting of a nucleic acid sequence selected from the group consisting of SEQ ID NO:78, SEQ ID NO:79, SEQ ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEQ ID NO:85, SEQ ID NO:86, SEQ ID NO:87, SEQ ID NO:88, SEQ ID NO:89, SEQ ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID NO:115, SEQ ID NO:116, SEQ ID NO:117, SEQ ID NO:118, SEQ ID NO:119, SEQ ID NO:120, SEQ ID NO:121, SEQ ID NO:122, SEQ ID NO:123, SEQ ID NO:124, SEQ ID NO:125, SEQ ID NO:126, SEQ ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:132, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139, SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154.

It will be appreciated by those skilled in the art that as a result of the degeneracy of the genetic code, a multitude of polynucleotide sequences encoding SIGP, some bearing minimal homology to the polynucleotide sequences of any known and naturally occurring gene, may be produced. Thus, the invention contemplates each and every possible variation of polynucleotide sequence that could be made by selecting combinations based on possible codon choices. These combinations are made in accordance with the standard triplet genetic code as applied to the polynucleotide sequence of naturally occurring SIGP, and all such

20

25

30

5

10

## PF-0459 US

variations are to be considered as being specifically disclosed.

Although nucleotide sequences which encode SIGP and its variants are preferably capable of hybridizing to the nucleotide sequence of the naturally occurring SIGP under appropriately selected conditions of stringency, it may be advantageous to produce nucleotide sequences encoding SIGP or its derivatives possessing a substantially different codon usage. Codons may be selected to increase the rate at which expression of the peptide occurs in a particular prokaryotic or eukaryotic host in accordance with the frequency with which particular codons are utilized by the host. Other reasons for substantially altering the nucleotide sequence encoding SIGP and its derivatives without altering the encoded amino acid sequences include the production of RNA transcripts having more desirable properties, such as a greater half-life, than transcripts produced from the naturally occurring sequence.

The invention also encompasses production of DNA sequences which encode SIGP and SIGP derivatives, or fragments thereof, entirely by synthetic chemistry. After production, the synthetic sequence may be inserted into any of the many available expression vectors and cell systems using reagents that are well known in the art. Moreover, synthetic chemistry may be used to introduce mutations into a sequence encoding SIGP or any fragment thereof.

Also encompassed by the invention are polynucleotide sequences that are capable of hybridizing to the claimed polynucleotide sequences, and, in particular, to those shown in SEQ ID NO:78, SEQ ID NO:79, SEQ ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEQ ID NO:85, SEQ ID NO:86, SEQ ID NO:87, SEQ ID NO:88, SEQ ID NO:89, SEQ ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID NO:115, SEQ ID NO:116, SEQ ID NO:117, SEQ ID NO:118, SEQ ID NO:119, SEQ ID NO:120, SEQ ID NO:121, SEQ ID NO:122, SEQ ID NO:123, SEQ ID NO:124, SEQ ID NO:125, SEQ ID NO:126, SEQ ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:131, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139,

20

25

30

5

10

PF-0459 US

SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154, under various conditions of stringency. (See, e.g., Wahl, G.M. and S.L. Berger (1987) Methods Enzymol. 152:399-407; and Kimmel, A.R. (1987) Methods Enzymol. 152:507-511.)

Methods for DNA sequencing are well known and generally available in the art and may be used to practice any of the embodiments of the invention. The methods may employ such enzymes as the Klenow fragment of DNA polymerase I, Sequenase® (US Biochemical Corp., Cleveland, OH), Taq polymerase (Perkin Elmer), thermostable T7 polymerase (Amersham, Chicago, IL), or combinations of polymerases and proofreading exonucleases such as those found in the ELONGASE Amplification System (GIBCO/BRL, Gaithersburg, MD). Preferably, the process is automated with machines such as the Hamilton Micro Lab 2200 (Hamilton, Reno, NV), Peltier Thermal Cycler (PTC200; MJ Research, Watertown, MA) and the ABI Catalyst and 373 and 377 DNA Sequencers (Perkin Elmer).

The nucleic acid sequences encoding SIGP may be extended utilizing a partial nucleotide sequence and employing various methods known in the art to detect upstream sequences, such as promoters and regulatory elements. For example, one method which may be employed, restriction-site PCR, uses universal primers to retrieve unknown sequence adjacent to a known locus. (See, e.g., Sarkar, G. (1993) PCR Methods Applic. 2:318-322.) In particular, genomic DNA is first amplified in the presence of a primer complementary to a linker sequence within the vector and a primer specific to the region predicted to encode the gene. The amplified sequences are then subjected to a second round of PCR with the same linker primer and another specific primer internal to the first one. Products of each round of PCR are transcribed with an appropriate RNA polymerase and sequenced using reverse transcriptase.

Inverse PCR may also be used to amplify or extend sequences using divergent primers based on a known region. (See, e.g., Triglia, T. et al. (1988) Nucleic Acids Res. 16:8186.) The primers may be designed using commercially available software such as OLIGO 4.06 Primer Analysis software (National Biosciences Inc., Plymouth, MN) or another appropriate program to be about 22 to 30 nucleotides in length, to have a GC content of about 50% or

25

30

5

10

#### PF-0459 US

more, and to anneal to the target sequence at temperatures of about 68°C to 72°C. The method uses several restriction enzymes to generate a suitable fragment in the known region of a gene. The fragment is then circularized by intramolecular ligation and used as a PCR template.

Another method which may be used is capture PCR, which involves PCR amplification of DNA fragments adjacent to a known sequence in human and yeast artificial chromosome DNA. (See, e.g., Lagerstrom, M. et al. (1991) PCR Methods Applic. 1:111-119.) In this method, multiple restriction enzyme digestions and ligations may be used to place an engineered double-stranded sequence into an unknown fragment of the DNA molecule before performing PCR. Other methods which may be used to retrieve unknown sequences are known in the art. (See, e.g., Parker, J.D. et al. (1991) Nucleic Acids Res. 19:3055-3060.) Additionally, one may use PCR, nested primers, and PromoterFinder™ libraries to walk genomic DNA (Clontech, Palo Alto, CA). This process avoids the need to screen libraries and is useful in finding intron/exon junctions.

When screening for full-length cDNAs, it is preferable to use libraries that have been size-selected to include larger cDNAs. Also, random-primed libraries are preferable in that they will include more sequences which contain the 5' regions of genes. Use of a randomly primed library may be especially preferable for situations in which an oligo d(T) library does not yield a full-length cDNA. Genomic libraries may be useful for extension of sequence into 5' non-transcribed regulatory regions.

Capillary electrophoresis systems which are commercially available may be used to analyze the size or confirm the nucleotide sequence of sequencing or PCR products. In particular, capillary sequencing may employ flowable polymers for electrophoretic separation, four different fluorescent dyes (one for each nucleotide) which are laser activated, and a charge coupled device camera for detection of the emitted wavelengths. Output/light intensity may be converted to electrical signal using appropriate software (e.g., Genotyper<sup>TM</sup> and Sequence Navigator<sup>TM</sup>, Perkin Elmer), and the entire process from loading of samples to computer analysis and electronic data display may be computer controlled. Capillary electrophoresis is especially preferable for the sequencing of small pieces of DNA which might be present in limited amounts in a particular sample.

10

15

20

25

30

## PF-0459 US

In another embodiment of the invention, polynucleotide sequences or fragments thereof which encode SIGP may be used in recombinant DNA molecules to direct expression of SIGP, or fragments or functional equivalents thereof, in appropriate host cells. Due to the inherent degeneracy of the genetic code, other DNA sequences which encode substantially the same or a functionally equivalent amino acid sequence may be produced, and these sequences may be used to clone and express SIGP.

As will be understood by those of skill in the art, it may be advantageous to produce SIGP-encoding nucleotide sequences possessing non-naturally occurring codons. For example, codons preferred by a particular prokaryotic or eukaryotic host can be selected to increase the rate of protein expression or to produce an RNA transcript having desirable properties, such as a half-life which is longer than that of a transcript generated from the naturally occurring sequence.

The nucleotide sequences of the present invention can be engineered using methods generally known in the art in order to alter SIGP-encoding sequences for a variety of reasons including, but not limited to, alterations which modify the cloning, processing, and/or expression of the gene product. DNA shuffling by random fragmentation and PCR reassembly of gene fragments and synthetic oligonucleotides may be used to engineer the nucleotide sequences. For example, site-directed mutagenesis may be used to insert new restriction sites, alter glycosylation patterns, change codon preference, produce splice variants, introduce mutations, and so forth.

In another embodiment of the invention, natural, modified, or recombinant nucleic acid sequences encoding SIGP may be ligated to a heterologous sequence to encode a fusion protein. For example, to screen peptide libraries for inhibitors of SIGP activity, it may be useful to encode a chimeric SIGP protein that can be recognized by a commercially available antibody. A fusion protein may also be engineered to contain a cleavage site located between the SIGP encoding sequence and the heterologous protein sequence, so that SIGP may be cleaved and purified away from the heterologous moiety.

In another embodiment, sequences encoding SIGP may be synthesized, in whole or in part, using chemical methods well known in the art. (See, e.g., Caruthers, M.H. et al. (1980) Nucl. Acids Res. Symp. Ser. 215-223, and Horn, T. et al. (1980) Nucl. Acids Res. Symp. Ser.

10

15

20

25

30

## PF-0459 US

225-232.) Alternatively, the protein itself may be produced using chemical methods to synthesize the amino acid sequence of SIGP, or a fragment thereof. For example, peptide synthesis can be performed using various solid-phase techniques. (See, e.g., Roberge, J.Y. et al. (1995) Science 269:202-204.) Automated synthesis may be achieved using the ABI 431A Peptide Synthesizer (Perkin Elmer).

The newly synthesized peptide may be substantially purified by preparative high performance liquid chromatography. (See, e.g, Chiez, R.M. and F.Z. Regnier (1990) Methods Enzymol. 182:392-421.) The composition of the synthetic peptides may be confirmed by amino acid analysis or by sequencing. (See, e.g., Creighton, T. (1983) Proteins, Structures and Molecular Properties, WH Freeman and Co., New York, NY.) Additionally, the amino acid sequence of SIGP, or any part thereof, may be altered during direct synthesis and/or combined with sequences from other proteins, or any part thereof, to produce a variant polypeptide.

In order to express a biologically active SIGP, the nucleotide sequences encoding SIGP or derivatives thereof may be inserted into appropriate expression vector, i.e., a vector which contains the necessary elements for the transcription and translation of the inserted coding sequence.

Methods which are well known to those skilled in the art may be used to construct expression vectors containing sequences encoding SIGP and appropriate transcriptional and translational control elements. These methods include <u>in vitro</u> recombinant DNA techniques, synthetic techniques, and <u>in vivo</u> genetic recombination. (See, e.g., Sambrook, J. et al. (1989) <u>Molecular Cloning, A Laboratory Manual</u>, Cold Spring Harbor Press, Plainview, NY, ch. 4, 8, and 16-17; and Ausubel, F.M. et al. (1995, and periodic supplements) <u>Current Protocols in Molecular Biology</u>, John Wiley & Sons, New York, NY, ch. 9, 13, and 16.)

A variety of expression vector/host systems may be utilized to contain and express sequences encoding SIGP. These include, but are not limited to, microorganisms such as bacteria transformed with recombinant bacteriophage, plasmid, or cosmid DNA expression vectors; yeast transformed with yeast expression vectors; insect cell systems infected with virus expression vectors (e.g., baculovirus); plant cell systems transformed with virus expression vectors (e.g., cauliflower mosaic virus (CaMV) or tobacco mosaic virus (TMV))

10

PF-0459 US

25

30

or with bacterial expression vectors (e.g., Ti or pBR322 plasmids); or animal cell systems. The invention is not limited by the host cell employed.

The "control elements" or "regulatory sequences" are those non-translated regions, e.g., enhancers, promoters, and 5' and 3' untranslated regions, of the vector and polynucleotide sequences encoding SIGP which interact with host cellular proteins to carry out transcription and translation. Such elements may vary in their strength and specificity. Depending on the vector system and host utilized, any number of suitable transcription and translation elements, including constitutive and inducible promoters, may be used. For example, when cloning in bacterial systems, inducible promoters, e.g., hybrid lacZ promoter of the Bluescript® phagemid (Stratagene, La Jolla, CA) or pSport1<sup>TM</sup> plasmid (GIBCO/BRL), may be used. The baculovirus polyhedrin promoter may be used in insect cells. Promoters or enhancers derived from the genomes of plant cells (e.g., heat shock, RUBISCO, and storage protein genes) or from plant viruses (e.g., viral promoters or leader sequences) may be cloned into the vector. In mammalian cell systems, promoters from mammalian genes or from mammalian viruses are preferable. If it is necessary to generate a cell line that contains multiple copies of the sequence encoding SIGP, vectors based on SV40 or EBV may be used with an appropriate selectable marker.

In bacterial systems, a number of expression vectors may be selected depending upon the use intended for SIGP. For example, when large quantities of SIGP are needed for the induction of antibodies, vectors which direct high level expression of fusion proteins that are readily purified may be used. Such vectors include, but are not limited to, multifunctional E. coli cloning and expression vectors such as Bluescript® (Stratagene), in which the sequence encoding SIGP may be ligated into the vector in frame with sequences for the amino-terminal Met and the subsequent 7 residues of \( \beta\)-galactosidase so that a hybrid protein is produced, and pIN vectors. (See, e.g., Van Heeke, G. and S.M. Schuster (1989) J. Biol. Chem. 264:5503-5509.) pGEX vectors (Pharmacia Biotech, Uppsala, Sweden) may also be used to express foreign polypeptides as fusion proteins with glutathione S-transferase (GST). In general, such fusion proteins are soluble and can easily be purified from lysed cells by adsorption to glutathione-agarose beads followed by elution in the presence of free glutathione. Proteins made in such systems may be designed to include heparin, thrombin, or

30

5

10

## PF-0459 US

factor XA protease cleavage sites so that the cloned polypeptide of interest can be released from the GST moiety at will.

In the yeast <u>Saccharomyces cerevisiae</u>, a number of vectors containing constitutive or inducible promoters, such as alpha factor, alcohol oxidase, and PGH, may be used. (See, e.g., Ausubel, <u>supra</u>; and Grant et al. (1987) Methods Enzymol. 153:516-544.)

In cases where plant expression vectors are used, the expression of sequences encoding SIGP may be driven by any of a number of promoters. For example, viral promoters such as the 35S and 19S promoters of CaMV may be used alone or in combination with the omega leader sequence from TMV. (Takamatsu, N. (1987) EMBO J. 6:307-311.) Alternatively, plant promoters such as the small subunit of RUBISCO or heat shock promoters may be used. (See, e.g., Coruzzi, G. et al. (1984) EMBO J. 3:1671-1680; Broglie, R. et al. (1984) Science 224:838-843; and Winter, J. et al. (1991) Results Probl. Cell Differ. 17:85-105.) These constructs can be introduced into plant cells by direct DNA transformation or pathogen-mediated transfection. Such techniques are described in a number of generally available reviews. (See, e.g., Hobbs, S. or Murry, L.E. in McGraw Hill Yearbook of Science and Technology (1992) McGraw Hill, New York, NY; pp. 191-196.)

An insect system may also be used to express SIGP. For example, in one such system, Autographa californica nuclear polyhedrosis virus (AcNPV) is used as a vector to express foreign genes in Spodoptera frugiperda cells or in Trichoplusia larvae. The sequences encoding SIGP may be cloned into a non-essential region of the virus, such as the polyhedrin gene, and placed under control of the polyhedrin promoter. Successful insertion of sequences encoding SIGP will render the polyhedrin gene inactive and produce recombinant virus lacking coat protein. The recombinant viruses may then be used to infect, for example, S. frugiperda cells or Trichoplusia larvae in which SIGP may be expressed. (See, e.g., Engelhard, E.K. et al. (1994) Proc. Nat. Acad. Sci. 91:3224-3227.)

In mammalian host cells, a number of viral-based expression systems may be utilized. In cases where an adenovirus is used as an expression vector, sequences encoding SIGP may be ligated into an adenovirus transcription/translation complex consisting of the late promoter and tripartite leader sequence. Insertion in a non-essential E1 or E3 region of the viral genome may be used to obtain a viable virus which is capable of expressing SIGP in infected

25

30

5

# PF-0459 US

host cells. (See, e.g., Logan, J. and T. Shenk (1984) Proc. Natl. Acad. Sci. 81:3655-3659.) In addition, transcription enhancers, such as the Rous sarcoma virus (RSV) enhancer, may be used to increase expression in mammalian host cells.

Human artificial chromosomes (HACs) may also be employed to deliver larger fragments of DNA than can be contained and expressed in a plasmid. HACs of about 6 kb to 10 Mb are constructed and delivered via conventional delivery methods (liposomes, polycationic amino polymers, or vesicles) for therapeutic purposes.

Specific initiation signals may also be used to achieve more efficient translation of sequences encoding SIGP. Such signals include the ATG initiation codon and adjacent sequences. In cases where sequences encoding SIGP and its initiation codon and upstream sequences are inserted into the appropriate expression vector, no additional transcriptional or translational control signals may be needed. However, in cases where only coding sequence, or a fragment thereof, is inserted, exogenous translational control signals including the ATG initiation codon should be provided. Furthermore, the initiation codon should be in the correct reading frame to ensure translation of the entire insert. Exogenous translational elements and initiation codons may be of various origins, both natural and synthetic. The efficiency of expression may be enhanced by the inclusion of enhancers appropriate for the particular cell system used. (See, e.g., Scharf, D. et al. (1994) Results Probl. Cell Differ. 20:125-162.)

In addition, a host cell strain may be chosen for its ability to modulate expression of the inserted sequences or to process the expressed protein in the desired fashion. Such modifications of the polypeptide include, but are not limited to, acetylation, carboxylation, glycosylation, phosphorylation, lipidation, and acylation. Post-translational processing which cleaves a "prepro" form of the protein may also be used to facilitate correct insertion, folding, and/or function. Different host cells which have specific cellular machinery and characteristic mechanisms for post-translational activities (e.g., CHO, HeLa, MDCK, HEK293, and WI38), are available from the American Type Culture Collection (ATCC, Bethesda, MD) and may be chosen to ensure the correct modification and processing of the foreign protein.

For long term, high yield production of recombinant proteins, stable expression is

30

5

10

## PF-0459 US

preferred. For example, cell lines capable of stably expressing SIGP can be transformed using expression vectors which may contain viral origins of replication and/or endogenous expression elements and a selectable marker gene on the same or on a separate vector. Following the introduction of the vector, cells may be allowed to grow for about 1 to 2 days in enriched media before being switched to selective media. The purpose of the selectable marker is to confer resistance to selection, and its presence allows growth and recovery of cells which successfully express the introduced sequences. Resistant clones of stably transformed cells may be proliferated using tissue culture techniques appropriate to the cell type.

Any number of selection systems may be used to recover transformed cell lines. These include, but are not limited to, the herpes simplex virus thymidine kinase genes and adenine phosphoribosyltransferase genes, which can be employed in tk or apr cells, respectively. (See, e.g., Wigler, M. et al. (1977) Cell 11:223-232; and Lowy, I. et al. (1980) Cell 22:817-823) Also, antimetabolite, antibiotic, or herbicide resistance can be used as the basis for selection. For example, dhfr confers resistance to methotrexate; npt confers resistance to the aminoglycosides neomycin and G-418; and als or pat confer resistance to chlorsulfuron and phosphinotricin acetyltransferase, respectively. (See, e.g., Wigler, M. et al. (1980) Proc. Natl. Acad. Sci. 77:3567-3570; Colbere-Garapin, F. et al (1981) J. Mol. Biol. 150:1-14; and Murry, supra.) Additional selectable genes have been described, e.g., trpB, which allows cells to utilize indole in place of tryptophan, or hisD, which allows cells to utilize histinol in place of histidine. (See, e.g., Hartman, S.C. and R.C. Mulligan (1988) Proc. Natl. Acad. Sci. 85:8047-8051.) Recently, the use of visible markers has gained popularity with such markers as anthocyanins, ß glucuronidase and its substrate GUS, luciferase and its substrate luciferin. Green fluorescent proteins (GFP) (Clontech, Palo Alto, CA) are also used (See, e.g., Chalfie, M. et al. (1994) Science 263:802-805.) These markers can be used not only to identify transformants, but also to quantify the amount of transient or stable protein expression attributable to a specific vector system. (See, e.g., Rhodes, C.A. et al. (1995) Methods Mol. Biol. 55:121-131.)

Although the presence/absence of marker gene expression suggests that the gene of interest is also present, the presence and expression of the gene may need to be confirmed.

10

15

20

25

30

#### PF-0459 US

For example, if the sequence encoding SIGP is inserted within a marker gene sequence, transformed cells containing sequences encoding SIGP can be identified by the absence of marker gene function. Alternatively, a marker gene can be placed in tandem with a sequence encoding SIGP under the control of a single promoter. Expression of the marker gene in response to induction or selection usually indicates expression of the tandem gene as well.

Alternatively, host cells which contain the nucleic acid sequence encoding SIGP and express SIGP may be identified by a variety of procedures known to those of skill in the art. These procedures include, but are not limited to, DNA-DNA or DNA-RNA hybridizations and protein bioassay or immunoassay techniques which include membrane, solution, or chip based technologies for the detection and/or quantification of nucleic acid or protein sequences.

The presence of polynucleotide sequences encoding SIGP can be detected by DNA-DNA or DNA-RNA hybridization or amplification using probes or fragments or fragments of polynucleotides encoding SIGP. Nucleic acid amplification based assays involve the use of oligonucleotides or oligomers based on the sequences encoding SIGP to detect transformants containing DNA or RNA encoding SIGP.

A variety of protocols for detecting and measuring the expression of SIGP, using either polyclonal or monoclonal antibodies specific for the protein, are known in the art. Examples of such techniques include enzyme-linked immunosorbent assays (ELISAs), radioimmunoassays (RIAs), and fluorescence activated cell sorting (FACS). A two-site, monoclonal-based immunoassay utilizing monoclonal antibodies reactive to two non-interfering epitopes on SIGP is preferred, but a competitive binding assay may be employed. These and other assays are well described in the art. (See, e.g., Hampton, R. et al. (1990) Serological Methods, a Laboratory Manual, APS Press, St Paul, MN, Section IV; and Maddox, D.E. et al. (1983) J. Exp. Med. 158:1211-1216).

A wide variety of labels and conjugation techniques are known by those skilled in the art and may be used in various nucleic acid and amino acid assays. Means for producing labeled hybridization or PCR probes for detecting sequences related to polynucleotides encoding SIGP include oligolabeling, nick translation, end-labeling, or PCR amplification using a labeled nucleotide. Alternatively, the sequences encoding SIGP, or any fragments thereof,

30

5

10

### PF-0459 US

may be cloned into a vector for the production of an mRNA probe. Such vectors are known in the art, are commercially available, and may be used to synthesize RNA probes <u>in vitro</u> by addition of an appropriate RNA polymerase such as T7, T3, or SP6 and labeled nucleotides. These procedures may be conducted using a variety of commercially available kits, such as those provided by Pharmacia & Upjohn (Kalamazoo, MI), Promega (Madison, WI), and U.S. Biochemical Corp. (Cleveland, OH). Suitable reporter molecules or labels which may be used for ease of detection include radionuclides, enzymes, fluorescent, chemiluminescent, or chromogenic agents, as well as substrates, cofactors, inhibitors, magnetic particles, and the like.

Host cells transformed with nucleotide sequences encoding SIGP may be cultured under conditions suitable for the expression and recovery of the protein from cell culture. The protein produced by a transformed cell may be secreted or contained intracellularly depending on the sequence and/or the vector used. As will be understood by those of skill in the art, expression vectors containing polynucleotides which encode SIGP may be designed to contain signal sequences which direct secretion of SIGP through a prokaryotic or eukaryotic cell membrane. Other constructions may be used to join sequences encoding SIGP to nucleotide sequences encoding a polypeptide domain which will facilitate purification of soluble proteins. Such purification facilitating domains include, but are not limited to, metal chelating peptides such as histidine-tryptophan modules that allow purification on immobilized metals, protein A domains that allow purification on immobilized immunoglobulin, and the domain utilized in the FLAGS extension/affinity purification system (Immunex Corp., Seattle, WA). The inclusion of cleavable linker sequences, such as those specific for Factor XA or enterokinase (Invitrogen, San Diego, CA), between the purification domain and the SIGP encoding sequence may be used to facilitate purification. One such expression vector provides for expression of a fusion protein containing SIGP and a nucleic acid encoding 6 histidine residues preceding a thioredoxin or an enterokinase cleavage site. The histidine residues facilitate purification on immobilized metal ion affinity chromatography. (IMAC) (See, e.g., Porath, J. et al. (1992) Prot. Exp. Purif. 3: 263-281.) The enterokinase cleavage site provides a means for purifying SIGP from the fusion protein. (See, e.g., Kroll, D.J. et al. (1993) DNA Cell Biol. 12:441-453.)

### PF-0459 US

Fragments of SIGP may be produced not only by recombinant production, but also by direct peptide synthesis using solid-phase techniques. (See, e.g., Creighton, T.E. (1984) Protein: Structures and Molecular Properties, pp. 55-60, W.H. Freeman and Co., New York, NY.) Protein synthesis may be performed by manual techniques or by automation.

Automated synthesis may be achieved, for example, using the Applied Biosystems 431A Peptide Synthesizer (Perkin Elmer). Various fragments of SIGP may be synthesized separately and then combined to produce the full length molecule.

## **THERAPEUTICS**

The expression of the human signal peptide-containing proteins of the invention (SIGP) is closely associated with cell proliferation. Therefore, in cancers or immune response where SIGP is an activator, transcription factor, or enhancer, and is promoting cell proliferation, it is desirable to decrease the expression of SIGP. In conditions where SIGP is an inhibitor or suppressor and is controlling or decreasing cell proliferation, it is desirable to provide the protein or to increase the expression of SIGP.

In one embodiment, where SIGP is an inhibitor, SIGP or a fragment or derivative thereof may be administered to a subject to treat or prevent a cancer such as adenocarcinoma, leukemia, lymphoma, melanoma, myeloma, sarcoma, and teratocarcinoma. Such cancers include, but are not limited to, cancers of the adrenal gland, bladder, bone, bone marrow, brain, breast, cervix, gall bladder, ganglia, gastrointestinal tract, heart, kidney, liver, lung, muscle, ovary, pancreas, parathyroid, penis, prostate, salivary glands, skin, spleen, testis, thymus, thyroid, and uterus.

In another embodiment, a pharmaceutical composition comprising purified SIGP may be used to treat or prevent a cancer including, but not limited to, those listed above.

In another embodiment, an agonist which is specific for SIGP may be administered to a subject to treat or prevent a cancer including, but not limited to, those cancers listed above.

In another further embodiment, a vector capable of expressing SIGP, or a fragment or a derivative thereof, may be administered to a subject to treat or prevent a cancer including, but not limited to, those cancers listed above.

In a further embodiment where SIGP is promoting cell proliferation, antagonists which

25

30

10

15

20

25

30

#### PF-0459 US

decrease the expression or activity of SIGP may be administered to a subject to treat or prevent a cancer such as adenocarcinoma, leukemia, lymphoma, melanoma, myeloma, sarcoma, and teratocarcinoma. Such cancers include, but are not limited to, cancers of the adrenal gland, bladder, bone, bone marrow, brain, breast, cervix, gall bladder, ganglia, gastrointestinal tract, heart, kidney, liver, lung, muscle, ovary, pancreas, parathyroid, penis, prostate, salivary glands, skin, spleen, testis, thymus, thyroid, and uterus. In one aspect, antibodies which specifically bind SIGP may be used directly as an antagonist or indirectly as a targeting or delivery mechanism for bringing a pharmaceutical agent to cells or tissue which express SIGP.

In another embodiment, a vector expressing the complement of the polynucleotide encoding SIGP may be administered to a subject to treat or prevent a cancer including, but not limited to, those cancers listed above.

In yet another embodiment where SIGP is promoting leukocyte activity or proliferation, antagonists which decrease the activity of SIGP may be administered to a subject to treat or prevent an immune response. Such responses include, but are not limited to, disorders such as AIDS, Addison's disease, adult respiratory distress syndrome, allergies, anemia, asthma, atherosclerosis, bronchitis, cholecystitus, Crohn's disease, ulcerative colitis, atopic dermatitis, dermatomyositis, diabetes mellitus, emphysema, atrophic gastritis, glomerulonephritis, gout, Graves' disease, hypereosinophilia, irritable bowel syndrome, lupus erythematosus, multiple sclerosis, myasthenia gravis, myocardial or pericardial inflammation, osteoarthritis, osteoporosis, pancreatitis, polymyositis, rheumatoid arthritis, scleroderma, Sjögren's syndrome, and autoimmune thyroiditis; complications of cancer, hemodialysis, extracorporeal circulation; viral, bacterial, fungal, parasitic, protozoal, and helminthic infections; and trauma. In one aspect, antibodies which specifically bind SIGP may be used directly as an antagonist or indirectly as a targeting or delivery mechanism for bringing a pharmaceutical agent to cells or tissue which express SIGP.

In another embodiment, a vector expressing the complement of the polynucleotide encoding SIGP may be administered to a subject to treat or prevent an immune response including, but not limited to, those listed above.

In other embodiments, any of the proteins, antagonists, antibodies, agonists,

30

5

10

#### PF-0459 US

complementary sequences, or vectors of the invention may be administered in combination with other appropriate therapeutic agents. Selection of the appropriate agents for use in combination therapy may be made by one of ordinary skill in the art, according to conventional pharmaceutical principles. The combination of therapeutic agents may act synergistically to effect the treatment or prevention of the various disorders described above. Using this approach, one may be able to achieve therapeutic efficacy with lower dosages of each agent, thus reducing the potential for adverse side effects.

An antagonist of SIGP may be produced using methods which are generally known in the art. In particular, purified SIGP may be used to produce antibodies or to screen libraries of pharmaceutical agents to identify those which specifically bind SIGP. Antibodies to SIGP may also be generated using methods that are well known in the art. Such antibodies may include, but are not limited to, polyclonal, monoclonal, chimeric, and single chain antibodies, Fab fragments, and fragments produced by a Fab expression library. Neutralizing antibodies (i.e., those which inhibit dimer formation) are especially preferred for therapeutic use.

For the production of antibodies, various hosts including goats, rabbits, rats, mice, humans, and others may be immunized by injection with SIGP or with any fragment or oligopeptide thereof which has immunogenic properties. Depending on the host species, various adjuvants may be used to increase immunological response. Such adjuvants include, but are not limited to, Freund's, mineral gels such as aluminum hydroxide, and surface active substances such as lysolecithin, pluronic polyols, polyanions, peptides, oil emulsions, KLH, and dinitrophenol. Among adjuvants used in humans, BCG (bacilli Calmette-Guerin) and Corvnebacterium parvum are especially preferable.

It is preferred that the oligopeptides, peptides, or fragments used to induce antibodies to SIGP have an amino acid sequence consisting of at least about 5 amino acids, and, more preferably, of at least about 10 amino acids. It is also preferable that these oligopeptides, peptides, or fragments are identical to a portion of the amino acid sequence of the natural protein and contain the entire amino acid sequence of a small, naturally occurring molecule. Short stretches of SIGP amino acids may be fused with those of another protein, such as KLH, and antibodies to the chimeric molecule may be produced.

Monoclonal antibodies to SIGP may be prepared using any technique which provides for

20

25

30

5

10

### PF-0459 US

the production of antibody molecules by continuous cell lines in culture. These include, but are not limited to, the hybridoma technique, the human B-cell hybridoma technique, and the EBV-hybridoma technique. (See, e.g., Kohler, G. et al. (1975) Nature 256:495-497; Kozbor, D. et al. (1985) J. Immunol. Methods 81:31-42; Cote, R.J. et al. (1983) Proc. Natl. Acad. Sci. 80:2026-2030; and Cole, S.P. et al. (1984) Mol. Cell Biol. 62:109-120.)

In addition, techniques developed for the production of "chimeric antibodies," such as the splicing of mouse antibody genes to human antibody genes to obtain a molecule with appropriate antigen specificity and biological activity, can be used. (See, e.g., Morrison, S.L. et al. (1984) Proc. Natl. Acad. Sci. 81:6851-6855; Neuberger, M.S. et al. (1984) Nature 312:604-608; and Takeda, S. et al. (1985) Nature 314:452-454.) Alternatively, techniques described for the production of single chain antibodies may be adapted, using methods known in the art, to produce SIGP-specific single chain antibodies. Antibodies with related specificity, but of distinct idiotypic composition, may be generated by chain shuffling from random combinatorial immunoglobulin libraries. (See, e.g., Burton D.R. (1991) Proc. Natl. Acad. Sci. 88:10134-10137.)

Antibodies may also be produced by inducing <u>in vivo</u> production in the lymphocyte population or by screening immunoglobulin libraries or panels of highly specific binding reagents as disclosed in the literature. (See, e.g., Orlandi, R. et al. (1989) Proc. Natl. Acad. Sci. 86: 3833-3837; and Winter, G. et al. (1991) Nature 349:293-299.)

Antibody fragments which contain specific binding sites for SIGP may also be generated. For example, such fragments include, but are not limited to, F(ab')2 fragments produced by pepsin digestion of the antibody molecule and Fab fragments generated by reducing the disulfide bridges of the F(ab')2 fragments. Alternatively, Fab expression libraries may be constructed to allow rapid and easy identification of monoclonal Fab fragments with the desired specificity. (See, e.g., Huse, W.D. et al. (1989) Science 246:1275-1281.)

Various immunoassays may be used for screening to identify antibodies having the desired specificity. Numerous protocols for competitive binding or immunoradiometric assays using either polyclonal or monoclonal antibodies with established specificities are well known in the art. Such immunoassays typically involve the measurement of complex

30

5

10

### PF-0459 US

formation between SIGP and its specific antibody. A two-site, monoclonal-based immunoassay utilizing monoclonal antibodies reactive to two non-interfering SIGP epitopes is preferred, but a competitive binding assay may also be employed. (Maddox, <u>supra</u>.)

In another embodiment of the invention, the polynucleotides encoding SIGP, or any fragment or complement thereof, may be used for therapeutic purposes. In one aspect, the complement of the polynucleotide encoding SIGP may be used in situations in which it would be desirable to block the transcription of the mRNA. In particular, cells may be transformed with sequences complementary to polynucleotides encoding SIGP. Thus, complementary molecules or fragments may be used to modulate SIGP activity, or to achieve regulation of gene function. Such technology is now well known in the art, and sense or antisense oligonucleotides or larger fragments can be designed from various locations along the coding or control regions of sequences encoding SIGP.

Expression vectors derived from retroviruses, adenoviruses, or herpes or vaccinia viruses, or from various bacterial plasmids, may be used for delivery of nucleotide sequences to the targeted organ, tissue, or cell population. Methods which are well known to those skilled in the art can be used to construct vectors which will express nucleic acid sequences complementary to the polynucleotides of the gene encoding SIGP. (See, e.g., Sambrook, supra; and Ausubel, supra.)

Genes encoding SIGP can be turned off by transforming a cell or tissue with expression vectors which express high levels of a polynucleotide, or fragment thereof, encoding SIGP. Such constructs may be used to introduce untranslatable sense or antisense sequences into a cell. Even in the absence of integration into the DNA, such vectors may continue to transcribe RNA molecules until they are disabled by endogenous nucleases. Transient expression may last for a month or more with a non-replicating vector, and may last even longer if appropriate replication elements are part of the vector system.

As mentioned above, modifications of gene expression can be obtained by designing complementary sequences or antisense molecules (DNA, RNA, or PNA) to the control, 5', or regulatory regions of the gene encoding SIGP. Oligonucleotides derived from the transcription initiation site, e.g., between about positions -10 and +10 from the start site, are preferred. Similarly, inhibition can be achieved using triple helix base-pairing methodology.

10

15

20

25

30

### PF-0459 US

Triple helix pairing is useful because it causes inhibition of the ability of the double helix to open sufficiently for the binding of polymerases, transcription factors, or regulatory molecules. Recent therapeutic advances using triplex DNA have been described in the literature. (See, e.g., Gee, J.E. et al. (1994) in Huber, B.E. and B.I. Carr, Molecular and Immunologic Approaches, Futura Publishing Co., Mt. Kisco, NY, pp. 163-177.) A complementary sequence or antisense molecule may also be designed to block translation of mRNA by preventing the transcript from binding to ribosomes.

Ribozymes, enzymatic RNA molecules, may also be used to catalyze the specific cleavage of RNA. The mechanism of ribozyme action involves sequence-specific hybridization of the ribozyme molecule to complementary target RNA, followed by endonucleolytic cleavage. For example, engineered hammerhead motif ribozyme molecules may specifically and efficiently catalyze endonucleolytic cleavage of sequences encoding SIGP.

Specific ribozyme cleavage sites within any potential RNA target are initially identified by scanning the target molecule for ribozyme cleavage sites, including the following sequences: GUA, GUU, and GUC. Once identified, short RNA sequences of between 15 and 20 ribonucleotides, corresponding to the region of the target gene containing the cleavage site, may be evaluated for secondary structural features which may render the oligonucleotide inoperable. The suitability of candidate targets may also be evaluated by testing accessibility to hybridization with complementary oligonucleotides using ribonuclease protection assays.

Complementary ribonucleic acid molecules and ribozymes of the invention may be prepared by any method known in the art for the synthesis of nucleic acid molecules. These include techniques for chemically synthesizing oligonucleotides such as solid phase phosphoramidite chemical synthesis. Alternatively, RNA molecules may be generated by in vitro and in vivo transcription of DNA sequences encoding SIGP. Such DNA sequences may be incorporated into a wide variety of vectors with suitable RNA polymerase promoters such as T7 or SP6. Alternatively, these cDNA constructs that synthesize complementary RNA, constitutively or inducibly, can be introduced into cell lines, cells, or tissues.

RNA molecules may be modified to increase intracellular stability and half-life.

Possible modifications include, but are not limited to, the addition of flanking sequences at

10 The state of th

25

30

5

the 5' and/or 3' ends of the molecule, or the use of phosphorothioate or 2' O-methyl rather than phosphodiesterase linkages within the backbone of the molecule. This concept is inherent in the production of PNAs and can be extended in all of these molecules by the inclusion of nontraditional bases such as inosine, queosine, and wybutosine, as well as acetylmethyl-, thio-, and similarly modified forms of adenine, cytidine, guanine, thymine, and uridine which are not as easily recognized by endogenous endonucleases.

Many methods for introducing vectors into cells or tissues are available and equally suitable for use in vivo, in vitro, and ex vivo. For ex vivo therapy, vectors may be introduced into stem cells taken from the patient and clonally propagated for autologous transplant back into that same patient. Delivery by transfection, by liposome injections, or by polycationic amino polymers may be achieved using methods which are well known in the art. (See, e.g., Goldman, C.K. et al. (1997) Nature Biotechnology 15:462-466.)

Any of the therapeutic methods described above may be applied to any subject in need of such therapy, including, for example, mammals such as dogs, cats, cows, horses, rabbits, monkeys, and most preferably, humans.

An additional embodiment of the invention relates to the administration of a pharmaceutical or sterile composition, in conjunction with a pharmaceutically acceptable carrier, for any of the therapeutic effects discussed above. Such pharmaceutical compositions may consist of SIGP, antibodies to SIGP, and mimetics, agonists, antagonists, or inhibitors of SIGP. The compositions may be administered alone or in combination with at least one other agent, such as a stabilizing compound, which may be administered in any sterile, biocompatible pharmaceutical carrier including, but not limited to, saline, buffered saline, dextrose, and water. The compositions may be administered to a patient alone, or in combination with other agents, drugs, or hormones.

The pharmaceutical compositions utilized in this invention may be administered by any number of routes including, but not limited to, oral, intravenous, intramuscular, intra-arterial, intramedullary, intrathecal, intraventricular, transdermal, subcutaneous, intraperitoneal, intranasal, enteral, topical, sublingual, or rectal means.

In addition to the active ingredients, these pharmaceutical compositions may contain suitable pharmaceutically-acceptable carriers comprising excipients and auxiliaries which

10

#### PF-0459 US

facilitate processing of the active compounds into preparations which can be used pharmaceutically. Further details on techniques for formulation and administration may be found in the latest edition of <u>Remington's Pharmaceutical Sciences</u> (Maack Publishing Co., Easton, PA).

Pharmaceutical compositions for oral administration can be formulated using pharmaceutically acceptable carriers well known in the art in dosages suitable for oral administration. Such carriers enable the pharmaceutical compositions to be formulated as tablets, pills, dragees, capsules, liquids, gels, syrups, slurries, suspensions, and the like, for ingestion by the patient.

Pharmaceutical preparations for oral use can be obtained through combining active compounds with solid excipient and processing the resultant mixture of granules (optionally, after grinding) to obtain tablets or dragee cores. Suitable auxiliaries can be added, if desired. Suitable excipients include carbohydrate or protein fillers, such as sugars, including lactose, sucrose, mannitol, and sorbitol; starch from corn, wheat, rice, potato, or other plants; cellulose, such as methyl cellulose, hydroxypropylmethyl-cellulose, or sodium carboxymethylcellulose; gums, including arabic and tragacanth; and proteins, such as gelatin and collagen. If desired, disintegrating or solubilizing agents may be added, such as the cross-linked polyvinyl pyrrolidone, agar, and alginic acid or a salt thereof, such as sodium alginate.

Dragee cores may be used in conjunction with suitable coatings, such as concentrated sugar solutions, which may also contain gum arabic, talc, polyvinylpyrrolidone, carbopol gel, polyethylene glycol, and/or titanium dioxide, lacquer solutions, and suitable organic solvents or solvent mixtures. Dyestuffs or pigments may be added to the tablets or dragee coatings for product identification or to characterize the quantity of active compound, i.e., dosage.

Pharmaceutical preparations which can be used orally include push-fit capsules made of gelatin, as well as soft, sealed capsules made of gelatin and a coating, such as glycerol or sorbitol. Push-fit capsules can contain active ingredients mixed with fillers or binders, such as lactose or starches, lubricants, such as talc or magnesium stearate, and, optionally, stabilizers. In soft capsules, the active compounds may be dissolved or suspended in suitable liquids, such as fatty oils, liquid, or liquid polyethylene glycol with or without stabilizers.

25

25

30

5

#### PF-0459 US

Pharmaceutical formulations suitable for parenteral administration may be formulated in aqueous solutions, preferably in physiologically compatible buffers such as Hanks's solution, Ringer's solution, or physiologically buffered saline. Aqueous injection suspensions may contain substances which increase the viscosity of the suspension, such as sodium carboxymethyl cellulose, sorbitol, or dextran. Additionally, suspensions of the active compounds may be prepared as appropriate oily injection suspensions. Suitable lipophilic solvents or vehicles include fatty oils, such as sesame oil, or synthetic fatty acid esters, such as ethyl oleate, triglycerides, or liposomes. Non-lipid polycationic amino polymers may also be used for delivery. Optionally, the suspension may also contain suitable stabilizers or agents to increase the solubility of the compounds and allow for the preparation of highly concentrated solutions.

For topical or nasal administration, penetrants appropriate to the particular barrier to be permeated are used in the formulation. Such penetrants are generally known in the art.

The pharmaceutical compositions of the present invention may be manufactured in a manner that is known in the art, e.g., by means of conventional mixing, dissolving, granulating, dragee-making, levigating, emulsifying, encapsulating, entrapping, or lyophilizing processes.

The pharmaceutical composition may be provided as a salt and can be formed with many acids, including but not limited to, hydrochloric, sulfuric, acetic, lactic, tartaric, malic, and succinic acid. Salts tend to be more soluble in aqueous or other protonic solvents than are the corresponding free base forms. In other cases, the preferred preparation may be a lyophilized powder which may contain any or all of the following: 1 mM to 50 mM histidine, 0.1% to 2% sucrose, and 2% to 7% mannitol, at a pH range of 4.5 to 5.5, that is combined with buffer prior to use.

After pharmaceutical compositions have been prepared, they can be placed in an appropriate container and labeled for treatment of an indicated condition. For administration of SIGP, such labeling would include amount, frequency, and method of administration.

Pharmaceutical compositions suitable for use in the invention include compositions wherein the active ingredients are contained in an effective amount to achieve the intended purpose. The determination of an effective dose is well within the capability of those skilled

5

PF-0459 US

in the art.

For any compound, the therapeutically effective dose can be estimated initially either in cell culture assays, e.g., of neoplastic cells or in animal models such as mice, rats, rabbits, dogs, or pigs. An animal model may also be used to determine the appropriate concentration range and route of administration. Such information can then be used to determine useful doses and routes for administration in humans.

A therapeutically effective dose refers to that amount of active ingredient, for example SIGP or fragments thereof, antibodies of SIGP, and agonists, antagonists or inhibitors of SIGP, which ameliorates the symptoms or condition. Therapeutic efficacy and toxicity may be determined by standard pharmaceutical procedures in cell cultures or with experimental animals, such as by calculating the ED50 (the dose therapeutically effective in 50% of the population) or LD50 (the dose lethal to 50% of the population) statistics. The dose ratio of therapeutic to toxic effects is the therapeutic index, and it can be expressed as the ED50/LD50 ratio. Pharmaceutical compositions which exhibit large therapeutic indices are preferred. The data obtained from cell culture assays and animal studies are used to formulate a range of dosage for human use. The dosage contained in such compositions is preferably within a range of circulating concentrations that includes the ED50 with little or no toxicity. The dosage varies within this range depending upon the dosage form employed, the sensitivity of the patient, and the route of administration.

The exact dosage will be determined by the practitioner, in light of factors related to the subject requiring treatment. Dosage and administration are adjusted to provide sufficient levels of the active moiety or to maintain the desired effect. Factors which may be taken into account include the severity of the disease state, the general health of the subject, the age, weight, and gender of the subject, time and frequency of administration, drug combination(s), reaction sensitivities, and response to therapy. Long-acting pharmaceutical compositions may be administered every 3 to 4 days, every week, or biweekly depending on the half-life and clearance rate of the particular formulation.

Normal dosage amounts may vary from about 0.1  $\mu$ g to 100,000  $\mu$ g, up to a total dose of about 1 gram, depending upon the route of administration. Guidance as to particular dosages and methods of delivery is provided in the literature and generally available to practitioners in

30

25

Ē

20

25

30

5

PF-0459 US

the art. Those skilled in the art will employ different formulations for nucleotides than for proteins or their inhibitors. Similarly, delivery of polynucleotides or polypeptides will be specific to particular cells, conditions, locations, etc.

## DIAGNOSTICS

In another embodiment, antibodies which specifically bind SIGP may be used for the diagnosis of disorders characterized by expression of SIGP, or in assays to monitor patients being treated with SIGP or agonists, antagonists, or inhibitors of SIGP. Antibodies useful for diagnostic purposes may be prepared in the same manner as described above for therapeutics. Diagnostic assays for SIGP include methods which utilize the antibody and a label to detect SIGP in human body fluids or in extracts of cells or tissues. The antibodies may be used with or without modification, and may be labeled by covalent or non-covalent attachment of a reporter molecule. A wide variety of reporter molecules, several of which are described above, are known in the art and may be used.

A variety of protocols for measuring SIGP, including ELISAs, RIAs, and FACS, are known in the art and provide a basis for diagnosing altered or abnormal levels of SIGP expression. Normal or standard values for SIGP expression are established by combining body fluids or cell extracts taken from normal mammalian subjects, preferably human, with antibody to SIGP under conditions suitable for complex formation. The amount of standard complex formation may be quantitated by various methods, preferably by photometric means. Quantities of SIGP expressed in subject, control, and disease samples from biopsied tissues are compared with the standard values. Deviation between standard and subject values establishes the parameters for diagnosing disease.

In another embodiment of the invention, the polynucleotides encoding SIGP may be used for diagnostic purposes. The polynucleotides which may be used include oligonucleotide sequences, complementary RNA and DNA molecules, and PNAs. The polynucleotides may be used to detect and quantitate gene expression in biopsied tissues in which expression of SIGP may be correlated with disease. The diagnostic assay may be used to determine absence, presence, and excess expression of SIGP, and to monitor regulation of SIGP levels during therapeutic intervention.

30

5

## PF-0459 US

In one aspect, hybridization with PCR probes which are capable of detecting polynucleotide sequences, including genomic sequences, encoding SIGP or closely related molecules may be used to identify nucleic acid sequences which encode SIGP. The specificity of the probe, whether it is made from a highly specific region, e.g., the 5' regulatory region, or from a less specific region, e.g., a conserved motif, and the stringency of the hybridization or amplification (maximal, high, intermediate, or low), will determine whether the probe identifies only naturally occurring sequences encoding SIGP, alleles, or related sequences.

Probes may also be used for the detection of related sequences, and should preferably contain at least 50% of the nucleotides from any of the SIGP encoding sequences. The hybridization probes of the subject invention may be DNA or RNA and may be derived from the sequence of SEQ ID NO:78, SEQ ID NO:79, SEQ ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEQ ID NO:85, SEQ ID NO:86, SEQ ID NO:87, SEQ ID NO:88, SEQ ID NO:89, SEQ ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID NO:115, SEQ ID NO:116, SEQ ID NO:117, SEQ ID NO:118, SEQ ID NO:119, SEQ ID NO:120, SEQ ID NO:121, SEQ ID NO:122, SEQ ID NO:123, SEQ ID NO:124, SEQ ID NO:125, SEQ ID NO:126, SEQ ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:132, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139, SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154, or from genomic sequences including promoters, enhancers, and introns of the SIGP gene.

Means for producing specific hybridization probes for DNAs encoding SIGP include the cloning of polynucleotide sequences encoding SIGP or SIGP derivatives into vectors for the

25

30

5

Ť0

production of mRNA probes. Such vectors are known in the art, are commercially available, and may be used to synthesize RNA probes <u>in vitro</u> by means of the addition of the appropriate RNA polymerases and the appropriate labeled nucleotides. Hybridization probes may be labeled by a variety of reporter groups, for example, by radionuclides such as <sup>32</sup>P or <sup>35</sup>S, or by enzymatic labels, such as alkaline phosphatase coupled to the probe via avidin/biotin coupling systems, and the like.

Polynucleotide sequences encoding SIGP may be used for the diagnosis of a disorder associated with either increased or decreased expression of SIGP. Examples of such a disorder include, but are not limited to, cancers such as adenocarcinoma, leukemia, lymphoma, melanoma, myeloma, sarcoma, teratocarcinoma, and cancers of the adrenal gland, bladder, bone, brain, breast, cervix, gall bladder, ganglia, gastrointestinal tract, heart, kidney, liver, lung, bone marrow, muscle, ovary, pancreas, parathyroid, penis, prostate, salivary glands, skin, spleen, testis, thymus, thyroid, and uterus; neuronal disorders such as akathesia, Alzheimer's disease, amnesia, amyotrophic lateral sclerosis, bipolar disorder, catatonia, cerebral neoplasms, dementia, depression, Down's syndrome, tardive dyskinesia, dystonias, epilepsy, Huntington's disease, multiple sclerosis, neurofibromatosis, Parkinson's disease, paranoid psychoses, schizophrenia, and Tourette's disorder; and immunological disorders such as AIDS, Addison's disease, adult respiratory distress syndrome, allergies, anemia, asthma, atherosclerosis, bronchitis, cholecystitus, Crohn's disease, ulcerative colitis, atopic dermatitis, dermatomyositis, diabetes mellitus, emphysema, atrophic gastritis, glomerulonephritis, gout, Graves' disease, hypereosinophilia, irritable bowel syndrome, lupus erythematosus, multiple sclerosis, myasthenia gravis, myocardial or pericardial inflammation, osteoarthritis, osteoporosis, pancreatitis, polymyositis, rheumatoid arthritis, scleroderma, Sjögren's syndrome, and thyroiditis. The polynucleotide sequences encoding SIGP may be used in Southern or northern analysis, dot blot, or other membrane-based technologies; in PCR technologies; in dipstick, pin, and ELISA assays; and in microarrays utilizing fluids or tissues from patients to detect altered SIGP expression. Such qualitative or quantitative methods are well known in the art.

In a particular aspect, the nucleotide sequences encoding SIGP may be useful in assays that detect the presence of associated disorders, particularly those mentioned above. The

25

30

5

#### PF-0459 US

nucleotide sequences encoding SIGP may be labeled by standard methods and added to a fluid or tissue sample from a patient under conditions suitable for the formation of hybridization complexes. After a suitable incubation period, the sample is washed and the signal is quantitated and compared with a standard value. If the amount of signal in the patient sample is significantly altered in comparison to a control sample then the presence of altered levels of nucleotide sequences encoding SIGP in the sample indicates the presence of the associated disorder. Such assays may also be used to evaluate the efficacy of a particular therapeutic treatment regimen in animal studies, in clinical trials, or to monitor the treatment of an individual patient.

In order to provide a basis for the diagnosis of a disorder associated with expression of SIGP, a normal or standard profile for expression is established. This may be accomplished by combining body fluids or cell extracts taken from normal subjects, either animal or human, with a sequence, or a fragment thereof, encoding SIGP, under conditions suitable for hybridization or amplification. Standard hybridization may be quantified by comparing the values obtained from normal subjects with values from an experiment in which a known amount of a substantially purified polynucleotide is used. Standard values obtained in this manner may be compared with values obtained from samples from patients who are symptomatic for a disorder. Deviation from standard values is used to establish the presence of a disorder.

Once the presence of a disorder is established and a treatment protocol is initiated, hybridization assays may be repeated on a regular basis to determine if the level of expression in the patient begins to approximate that which is observed in the normal subject. The results obtained from successive assays may be used to show the efficacy of treatment over a period ranging from several days to months.

With respect to cancer, the presence of a relatively high amount of transcript in biopsied tissue from an individual may indicate a predisposition for the development of the disease, or may provide a means for detecting the disease prior to the appearance of actual clinical symptoms. A more definitive diagnosis of this type may allow health professionals to employ preventative measures or aggressive treatment earlier thereby preventing the development or further progression of the cancer.

25

30

5

### PF-0459 US

Additional diagnostic uses for oligonucleotides designed from the sequences encoding SIGP may involve the use of PCR. These oligomers may be chemically synthesized, generated enzymatically, or produced <u>in vitro</u>. Oligomers will preferably contain a fragment of a polynucleotide encoding SIGP, or a fragment of a polynucleotide complementary to the polynucleotide encoding SIGP, and will be employed under optimized conditions for identification of a specific gene or condition. Oligomers may also be employed under less stringent conditions for detection or quantitation of closely related DNA or RNA sequences.

Methods which may also be used to quantitate the expression of SIGP include radiolabeling or biotinylating nucleotides, coamplification of a control nucleic acid, and interpolating results from standard curves. (See, e.g., Melby, P.C. et al. (1993) J. Immunol. Methods 159:235-244; and Duplaa, C. et al. (1993) Anal. Biochem. 229-236.) The speed of quantitation of multiple samples may be accelerated by running the assay in an ELISA format where the oligomer of interest is presented in various dilutions and a spectrophotometric or colorimetric response gives rapid quantitation.

In further embodiments, oligonucleotides or longer fragments derived from any of the polynucleotide sequences described herein may be used as targets in a microarray. The microarray can be used to monitor the expression level of large numbers of genes simultaneously and to identify genetic variants, mutations, and polymorphisms. This information may be used to determine gene function, to understand the genetic basis of a disorder, to diagnose a disorder, and to develop and monitor the activities of therapeutic agents.

In one embodiment, the microarray is prepared and used according to methods known in the art. (See, e.g., Chee et al. (1995) PCT application WO95/11995; Lockhart, D. J. et al. (1996) Nat. Biotech. 14:1675-1680; and Schena, M. et al. (1996) Proc. Natl. Acad. Sci. 93:10614-10619.)

The microarray is preferably composed of a large number of unique single-stranded nucleic acid sequences, usually either synthetic antisense oligonucleotides or fragments of cDNAs. The oligonucleotides are preferably about 6 to 60 nucleotides in length, more preferably about 15 to 30 nucleotides in length, and most preferably about 20 to 25 nucleotides in length. It may be preferable to use oligonucleotides which are about 7 to 10

20

25

30

10

## PF-0459 US

nucleotides in length. The microarray may contain oligonucleotides which cover the known 5' or 3' sequence, sequential oligonucleotides which cover the full length sequence, or unique oligonucleotides selected from particular areas along the length of the sequence.

Polynucleotides used in the microarray may be oligonucleotides specific to a gene or genes of interest. Oligonucleotides can also be specific to one or more unidentified cDNAs associated with a particular cell type or tissue type. It may be appropriate to use pairs of oligonucleotides on a microarray. The first oligonucleotide in each pair differs from the second oligonucleotide by one nucleotide. This nucleotide is preferably located in the center of the sequence. The second oligonucleotide serves as a control. The number of oligonucleotide pairs may range from about 2 to 1,000,000.

In order to produce oligonucleotides for use on a microarray, the gene of interest is examined using a computer algorithm which starts at the 5' end, or, more preferably, at the 3' end of the nucleotide sequence. The algorithm identifies oligomers of defined length that are unique to the gene, have a GC content within a range suitable for hybridization, and lack secondary structure that may interfere with hybridization. In one aspect, the oligomers may be synthesized on a substrate using a light-directed chemical process. (See, e.g., Chee et al., <a href="supra">supra</a>.) The substrate may be any suitable solid support, e.g., paper, nylon, any other type of membrane, or a filter, chip, or glass slide.

In another aspect, the oligonucleotides may be synthesized on the surface of the substrate using a chemical coupling procedure and an ink jet application apparatus. (See, e.g., Baldeschweiler et al. (1995) PCT application WO95/251116.) An array analogous to a dot or slot blot (HYBRIDOT® apparatus, GIBCO/BRL) may be used to arrange and link cDNA fragments or oligonucleotides to the surface of a substrate using a vacuum system or thermal, UV, mechanical, or chemical bonding procedures. An array may also be produced by hand or by using available devices, materials, and machines, e.g. Brinkmann® multichannel pipettors or robotic instruments. The array may contain from 2 to 1,000,000 or any other feasible number of oligonucleotides.

In order to conduct sample analysis using the microarrays, polynucleotides are extracted from a sample. The sample may be obtained from any bodily fluid, e.g., blood, urine, saliva, phlegm, gastric juices, cultured cells, biopsies, or other tissue preparations. To produce

30

5

probes, the polynucleotides extracted from the sample are used to produce nucleic acid sequences complementary to the nucleic acids on the microarray. If the microarray contains cDNAs, antisense RNAs (aRNAs) are appropriate probes. Therefore, in one aspect, mRNA is reverse-transcribed to cDNA. The cDNA, in the presence of fluorescent label, is used to produce fragment or oligonucleotide aRNA probes. The fluorescently labeled probes are incubated with the microarray so that the probes hybridize to the microarray oligonucleotides. Nucleic acid sequences used as probes can include polynucleotides, fragments, and complementary or antisense sequences produced using restriction enzymes, PCR, or other methods known in the art.

Hybridization conditions can be adjusted so that hybridization occurs with varying degrees of complementarity. A scanner can be used to determine the levels and patterns of fluorescence after removal of any nonhybridized probes. The degree of complementarity and the relative abundance of each oligonucleotide sequence on the microarray can be assessed through analysis of the scanned images. A detection system may be used to measure the absence, presence, or level of hybridization for any of the sequences. (See, e.g., Heller, R.A. et al. (1997) Proc. Natl. Acad. Sci. 94:2150-2155.)

In another embodiment of the invention, nucleic acid sequences encoding SIGP may be used to generate hybridization probes useful in mapping the naturally occurring genomic sequence. The sequences may be mapped to a particular chromosome, to a specific region of a chromosome, or to artificial chromosome constructions, e.g., human artificial chromosomes (HACs), yeast artificial chromosomes (YACs), bacterial artificial chromosomes (BACs), bacterial P1 constructions, or single chromosome cDNA libraries. (See, e.g., Price, C.M. (1993) Blood Rev. 7:127-134; and Trask, B.J. (1991) Trends Genet. 7:149-154.)

Fluorescent <u>in situ</u> hybridization (FISH) may be correlated with other physical chromosome mapping techniques and genetic map data. (See, e.g., Heinz-Ulrich, et al. (1995) in Meyers, R.A. (ed.) <u>Molecular Biology and Biotechnology</u>, VCH Publishers New York, NY, pp. 965-968.) Examples of genetic map data can be found in various scientific journals or at the Online Mendelian Inheritance in Man (OMIM) site. Correlation between the location of the gene encoding SIGP on a physical chromosomal map and a specific disorder, or a predisposition to a specific disorder, may help define the region of DNA

25

30

4

5

#### PF-0459 US

associated with that disorder. The nucleotide sequences of the invention may be used to detect differences in gene sequences among normal, carrier, and affected individuals.

In situ hybridization of chromosomal preparations and physical mapping techniques, such as linkage analysis using established chromosomal markers, may be used for extending genetic maps. Often the placement of a gene on the chromosome of another mammalian species, such as mouse, may reveal associated markers even if the number or arm of a particular human chromosome is not known. New sequences can be assigned to chromosomal arms by physical mapping. This provides valuable information to investigators searching for disease genes using positional cloning or other gene discovery techniques. Once the disease or syndrome has been crudely localized by genetic linkage to a particular genomic region, e.g., AT to 11q22-23, any sequences mapping to that area may represent associated or regulatory genes for further investigation. (See, e.g., Gatti, R.A. et al. (1988) Nature 336:577-580.) The nucleotide sequence of the subject invention may also be used to detect differences in the chromosomal location due to translocation, inversion, etc., among normal, carrier, or affected individuals.

In another embodiment of the invention, SIGP, its catalytic or immunogenic fragments, or oligopeptides thereof can be used for screening libraries of compounds in any of a variety of drug screening techniques. The fragment employed in such screening may be free in solution, affixed to a solid support, borne on a cell surface, or located intracellularly. The formation of binding complexes between SIGP and the agent being tested may be measured.

Another technique for drug screening provides for high throughput screening of compounds having suitable binding affinity to the protein of interest. (See, e.g., Geysen, et al. (1984) PCT application WO84/03564.) In this method, large numbers of different small test compounds are synthesized on a solid substrate, such as plastic pins or some other surface. The test compounds are reacted with SIGP, or fragments thereof, and washed. Bound SIGP is then detected by methods well known in the art. Purified SIGP can also be coated directly onto plates for use in the aforementioned drug screening techniques. Alternatively, non-neutralizing antibodies can be used to capture the peptide and immobilize it on a solid support.

In another embodiment, one may use competitive drug screening assays in which

30

5

10

neutralizing antibodies capable of binding SIGP specifically compete with a test compound for binding SIGP. In this manner, antibodies can be used to detect the presence of any peptide which shares one or more antigenic determinants with SIGP.

In additional embodiments, the nucleotide sequences which encode SIGP may be used in any molecular biology techniques that have yet to be developed, provided the new techniques rely on properties of nucleotide sequences that are currently known, including, but not limited to, such properties as the triplet genetic code and specific base pair interactions.

The examples below are provided to illustrate the subject invention and are not included for the purpose of limiting the invention.

### **EXAMPLES**

For purposes of example, the preparation and sequencing of the SPLNNOT04 cDNA library, from which Incyte Clones 1534876 and 1559131 were isolated, is described. Preparation and sequencing of cDNAs in libraries in the LIFESEQ<sup>TM</sup> database have varied over time, and the gradual changes involved use of kits, plasmids, and machinery available at the particular time the library was made and analyzed.

# I. SPLNNOT04 cDNA Library Construction

The SPLNNOT04 cDNA library was constructed from microscopically normal spleen tissue obtained from a 2-year-old Hispanic male who died of cerebral anoxia. The patient's serologies and past medical history were negative.

The frozen tissue was homogenized and lysed using a Brinkmann Homogenizer Polytron PT-3000 (Brinkmann Instruments, Westbury, NJ) in guanidinium isothiocyanate solution. The lysate was centrifuged over a 5.7 M CsCl cushion using an Beckman SW28 rotor in a Beckman L8-70M Ultracentrifuge (Beckman Instruments) for 18 hours at 25,000 rpm at ambient temperature. The RNA was extracted with acid phenol pH 4.0, precipitated using 0.3 M sodium acetate and 2.5 volumes of ethanol, resuspended in RNAse-free water and DNase treated at 37°C. The RNA extraction and precipitation were repeated as before. The mRNA was then isolated using the Qiagen Oligotex kit (QIAGEN Inc., Chatsworth, CA) and used to construct the cDNA library.

30

5

10

The mRNA was handled according to the recommended protocols in the SuperScript plasmid system (Cat. #18248-013, Gibco-BRL, Gaithersburg, MD). cDNA synthesis was initiated with a NotI-oligo d(T) primer. Double-stranded cDNA was blunted, ligated to EcoRI adaptors, digested with NotI, fractionated on a Sepharose CL4B column (Cat. #275105-01, Pharmacia), and those cDNAs exceeding 400 bp were ligated into the NotI and EcoRI sites of the pINCY 1 vector (Incyte). The plasmid pINCY 1 was subsequently transformed into DH5α<sup>TM</sup> competent cells (Cat. #18258-012, Gibco-BRL).

# II Isolation and Sequencing of cDNA Clones

Plasmid cDNA was released from the cells and purified using the REAL Prep 96 plasmid kit (Catalog #26173, QIAGEN). The recommended protocol was employed except for the following changes: 1) the bacteria were cultured in 1 ml of sterile Terrific Broth (Catalog #22711, GIBCO-BRL) with carbenicillin at 25 mg/L and glycerol at 0.4%; 2) after inoculation, the cultures were incubated for 19 hours and at the end of incubation, the cells were lysed with 0.3 ml of lysis buffer; and 3) following isopropanol precipitation, the plasmid DNA pellet was resuspended in 0.1 ml of distilled water. After the last step in the protocol, samples were transferred to a 96-well block for storage at 4° C.

cDNAs were sequenced according to the method of Sanger et al. (1975, J. Mol. Biol. 94:441f), using the Perkin Elmer Catalyst 800 or a Hamilton Micro Lab 2200 (Hamilton, Reno, NV) in combination with Peltier Thermal Cyclers (PTC200 from MJ Research, Watertown, MA) and Applied Biosystems 377 DNA Sequencing Systems or the Perkin Elmer 373 DNA Sequencing System and the reading frame was determined.

# III. Homology Searching of cDNA Clones and Their Deduced Proteins

The nucleotide sequences and/or amino acid sequences of the Sequence Listing were used to query sequences in the GenBank, SwissProt, BLOCKS, and Pima II databases. These databases, which contain previously identified and annotated sequences, were searched for regions of homology using BLAST (Basic Local Alignment Search Tool). (See, e.g., Altschul, S.F. (1993) J. Mol. Evol 36:290-300; and Altschul et al. (1990) J. Mol. Biol. 215:403-410.)

109

10 for the first the first

5

BLAST produced alignments of both nucleotide and amino acid sequences to determine sequence similarity. Because of the local nature of the alignments, BLAST was especially useful in determining exact matches or in identifying homologs which may be of prokaryotic (bacterial) or eukaryotic (animal, fungal, or plant) origin. Other algorithms could have been used when dealing with primary sequence patterns and secondary structure gap penalties. (See, e.g., Smith, T. et al. (1992) Protein Engineering 5:35-51.) The sequences disclosed in this application have lengths of at least 49 nucleotides and have no more than 12% uncalled bases (where N is recorded rather than A, C, G, or T).

The BLAST approach searched for matches between a query sequence and a database sequence. BLAST evaluated the statistical significance of any matches found, and reported only those matches that satisfy the user-selected threshold of significance. In this application, threshold was set at  $10^{-25}$  for nucleotides and  $10^{-8}$  for peptides.

Incyte nucleotide sequences were searched against the GenBank databases for primate (pri), rodent (rod), and other mammalian sequences (mam), and deduced amino acid sequences from the same clones were then searched against GenBank functional protein databases, mammalian (mamp), vertebrate (vrtp), and eukaryote (eukp), for homology.

# IV. Northern Analysis

Northern analysis is a laboratory technique used to detect the presence of a transcript of a gene and involves the hybridization of a labeled nucleotide sequence to a membrane on which RNAs from a particular cell type or tissue have been bound. (See, e.g., Sambrook, <u>supra</u>, ch. 7; and Ausubel, F.M. et al. <u>supra</u>, ch. 4 and 16.)

Analogous computer techniques applying BLAST are used to search for identical or related molecules in nucleotide databases such as GenBank or LIFESEQ™ database (Incyte Pharmaceuticals). This analysis is much faster than multiple membrane-based hybridizations. In addition, the sensitivity of the computer search can be modified to determine whether any particular match is categorized as exact or homologous.

The basis of the search is the product score, which is defined as:

% sequence identity x % maximum BLAST score

25

25

30

5

10

#### PF-0459 US

The product score takes into account both the degree of similarity between two sequences and the length of the sequence match. For example, with a product score of 40, the match will be exact within a 1% to 2% error, and, with a product score of 70, the match will be exact. Homologous molecules are usually identified by selecting those which show product scores between 15 and 40, although lower scores may identify related molecules.

The results of northern analysis are reported as a list of libraries in which the transcript encoding SIGP occurs. Abundance and percent abundance are also reported. Abundance directly reflects the number of times a particular transcript is represented in a cDNA library, and percent abundance is abundance divided by the total number of sequences examined in the cDNA library.

# V. Extension of SIGP Encoding Polynucleotides

The nucleic acid sequence of one of the polynucleotides of the present invention was used to design oligonucleotide primers for extending a partial nucleotide sequence to full length. One primer was synthesized to initiate extension of an antisense polynucleotide, and the other was synthesized to initiate extension of a sense polynucleotide. Primers were used to facilitate the extension of the known sequence "outward" generating amplicons containing new unknown nucleotide sequence for the region of interest. The initial primers were designed from the cDNA using OLIGO 4.06 (National Biosciences, Plymouth, MN), or another appropriate program, to be about 22 to 30 nucleotides in length, to have a GC content of about 50% or more, and to anneal to the target sequence at temperatures of about 68 °C to about 72 °C. Any stretch of nucleotides which would result in hairpin structures and primer-primer dimerizations was avoided.

Selected human cDNA libraries (GIBCO/BRL) were used to extend the sequence. If more than one extension is necessary or desired, additional sets of primers are designed to further extend the known region.

High fidelity amplification was obtained by following the instructions for the XL-PCR kit (Perkin Elmer) and thoroughly mixing the enzyme and reaction mix. PCR was performed using the Peltier Thermal Cycler (PTC200; M.J. Research, Watertown, MA), beginning with 40 pmol of each primer and the recommended concentrations of all other components of the

35

5

10

#### PF-0459 US

kit, with the following parameters:

	Step 1	94° C for 1 min (initial denaturation)
	Step 2	65° C for 1 min
	Step 3	68° C for 6 min
	Step 4	94° C for 15 sec
	Step 5	65° C for 1 min
	Step 6	68° C for 7 min
	Step 7	Repeat steps 4 through 6 for an additional 15 cycles
	Step 8	94° C for 15 sec
1	Step 9	65° C for 1 min
	Step 10	68° C for 7:15 min
	Step 11	Repeat steps 8 through 10 for an additional 12 cycles
	Step 12	72° C for 8 min
	Step 13	4° C (and holding)

A 5  $\mu$ l to 10  $\mu$ l aliquot of the reaction mixture was analyzed by electrophoresis on a low concentration (about 0.6% to 0.8%) agarose mini-gel to determine which reactions were successful in extending the sequence. Bands thought to contain the largest products were excised from the gel, purified using QIAQuick<sup>TM</sup> (QIAGEN Inc., Chatsworth, CA), and trimmed of overhangs using Klenow enzyme to facilitate religation and cloning.

After ethanol precipitation, the products were redissolved in 13  $\mu$ l of ligation buffer,  $1\mu$ l T4-DNA ligase (15 units) and  $1\mu$ l T4 polynucleotide kinase were added, and the mixture was incubated at room temperature for 2 to 3 hours, or overnight at 16° C. Competent E. coli cells (in 40  $\mu$ l of appropriate media) were transformed with 3  $\mu$ l of ligation mixture and cultured in 80  $\mu$ l of SOC medium. (See, e.g., Sambrook, supra, Appendix A, p. 2.) After incubation for one hour at 37° C, the E. coli mixture was plated on Luria Bertani (LB) agar (See, e.g., Sambrook, supra, Appendix A, p. 1) containing 2x Carb. The following day, several colonies were randomly picked from each plate and cultured in 150  $\mu$ l of liquid LB/2x Carb medium placed in an individual well of an appropriate commercially-available sterile 96-well microtiter plate. The following day, 5  $\mu$ l of each overnight culture was transferred into a non-sterile 96-well plate and, after dilution 1:10 with water, 5  $\mu$ l from each sample was transferred into a PCR array.

For PCR amplification, 18  $\mu$ l of concentrated PCR reaction mix (3.3x) containing 4 units of rTth DNA polymerase, a vector primer, and one or both of the gene specific primers used for the extension reaction were added to each well. Amplification was performed using

30

5

10

### PF-0459 US

the following conditions:

S	Step 1	94° C for 60 sec
S	Step 2	94° C for 20 sec
S	Step 3	55° C for 30 sec
S	Step 4	72° C for 90 sec
S	Step 5	Repeat steps 2 through 4 for an additional 29 cycles
S	Step 6	72° C for 180 sec
S	Step 7	4° C (and holding)

Aliquots of the PCR reactions were run on agarose gels together with molecular weight markers. The sizes of the PCR products were compared to the original partial cDNAs, and appropriate clones were selected, ligated into plasmid, and sequenced.

In like manner, the nucleotide sequence of one of the nucleotide sequences of the present invention were used to obtain 5' regulatory sequences using the procedure above, oligonucleotides designed for 5' extension, and an appropriate genomic library.

# VI. Labeling and Use of Individual Hybridization Probes

Hybridization probes derived from one of the nucleotide sequences of the present invention are employed to screen cDNAs, genomic DNAs, or mRNAs. Although the labeling of oligonucleotides, consisting of about 20 base pairs, is specifically described, essentially the same procedure is used with larger nucleotide fragments. Oligonucleotides are designed using state-of-the-art software such as OLIGO 4.06 (National Biosciences) and labeled by combining 50 pmol of each oligomer, 250  $\mu$ Ci of [ $\gamma$ - $^{32}$ P] adenosine triphosphate (Amersham, Chicago, IL), and T4 polynucleotide kinase (DuPont NEN®, Boston, MA). The labeled oligonucleotides are substantially purified using a Sephadex G-25 superfine resin column (Pharmacia & Upjohn, Kalamazoo, MI). An aliquot containing  $10^7$  counts per minute of the labeled probe is used in a typical membrane-based hybridization analysis of human genomic DNA digested with one of the following endonucleases: Ase I, Bgl II, Eco RI, Pst I, Xba 1, or Pvu II (DuPont NEN, Boston, MA).

The DNA from each digest is fractionated on a 0.7 percent agarose gel and transferred to nylon membranes (Nytran Plus, Schleicher & Schuell, Durham, NH). Hybridization is carried out for 16 hours at 40°C. To remove nonspecific signals, blots are sequentially washed at room temperature under increasingly stringent conditions up to 0.1 x

30

10

### PF-0459 US

saline sodium citrate and 0.5% sodium dodecyl sulfate. After XOMAT AR<sup>TM</sup> film (Kodak, Rochester, NY) is exposed to the blots to film for several hours, hybridization patterns are compared visually.

# 5 VII. Microarrays

To produce oligonucleotides for a microarray, one of the nucleotide sequences of the present invention is examined using a computer algorithm which starts at the 3' end of the nucleotide sequence. For each, the algorithm identifies oligomers of defined length that are unique to the nucleic acid sequence, have a GC content within a range suitable for hybridization, and lack secondary structure that would interfere with hybridization. The algorithm identifies approximately 20 oligonucleotides corresponding to each nucleic acid sequence. For each sequence-specific oligonucleotide, a pair of oligonucleotides is synthesized in which the first oligonucleotides differs from the second oligonucleotide by one nucleotide in the center of the sequence. The oligonucleotide pairs can be arranged on a substrate, e.g. a silicon chip, using a light-directed chemical process. (See, e.g., Chee, supra.)

In the alternative, a chemical coupling procedure and an ink jet device can be used to synthesize oligomers on the surface of a substrate. (See, e.g., Baldeschweiler, supra.) An array analogous to a dot or slot blot may also be used to arrange and link fragments or oligonucleotides to the surface of a substrate using or thermal, UV, mechanical, or chemical bonding procedures, or a vacuum system. A typical array may be produced by hand or using available methods and machines and contain any appropriate number of elements. After hybridization, nonhybridized probes are removed and a scanner used to determine the levels and patterns of fluorescence. The degree of complementarity and the relative abundance of each oligonucleotide sequence on the microarray may be assessed through analysis of the scanned images.

# VIII. Complementary Polynucleotides

Sequences complementary to the SIGP-encoding sequences, or any parts thereof, are used to detect, decrease, or inhibit expression of naturally occurring SIGP. Although use of oligonucleotides comprising from about 15 to 30 base pairs is described, essentially the same

10

| Compared to the compared

25

30

5

procedure is used with smaller or with larger sequence fragments. Appropriate oligonucleotides are designed using Oligo 4.06 software and the coding sequence of SIGP. To inhibit transcription, a complementary oligonucleotide is designed from the most unique 5' sequence and used to prevent promoter binding to the coding sequence. To inhibit translation, a complementary oligonucleotide is designed to prevent ribosomal binding to the SIGP-encoding transcript.

# IX. Expression of SIGP

Expression of SIGP is accomplished by subcloning the cDNA into an appropriate vector and transforming the vector into host cells. This vector contains an appropriate promoter, e.g., β-galactosidase upstream of the cloning site, operably associated with the cDNA of interest. (See, e.g.,Sambrook, supra, pp. 404-433; and Rosenberg, M. et al. (1983) Methods Enzymol. 101:123-138.)

Induction of an isolated, transformed bacterial strain with isopropyl beta-D-thiogalactopyranoside (IPTG) using standard methods produces a fusion protein which consists of the first 8 residues of \( \mathbb{B}\)-galactosidase, about 5 to 15 residues of linker, and the full length protein. The signal residues direct the secretion of SIGP into bacterial growth media which can be used directly in the following assay for activity.

# X. Production of SIGP Specific Antibodies

SIGP substantially purified using PAGE electrophoresis (see, e.g., Harrington, M.G. (1990) Methods Enzymol. 182:488-495), or other purification techniques, is used to immunize rabbits and to produce antibodies using standard protocols. The SIGP amino acid sequence is analyzed using DNASTAR software (DNASTAR Inc) to determine regions of high immunogenicity, and a corresponding oligopeptide is synthesized and used to raise antibodies by means known to those of skill in the art. Methods for selection of appropriate epitopes, such as those near the C-terminus or in hydrophilic regions are well described in the art. (See, e.g., Ausubel et al. supra, ch. 11.)

Typically, the oligopeptides are 15 residues in length, and are synthesized using an Applied Biosystems Peptide Synthesizer Model 431A using fmoc-chemistry and coupled to

30

5

10

KLH (Sigma, St. Louis, MO) by reaction with N-maleimidobenzoyl-N-hydroxysuccinimide ester (MBS) to increase immunogenicity. (See, e.g., Ausubel et al. <u>supra.</u>) Rabbits are immunized with the oligopeptide-KLH complex in complete Freund's adjuvant. Resulting antisera are tested for antipeptide activity, for example, by binding the peptide to plastic, blocking with 1% BSA, reacting with rabbit antisera, washing, and reacting with radio-iodinated goat anti-rabbit IgG.

# XI. Purification of Naturally Occurring SIGP Using Specific Antibodies

Naturally occurring or recombinant SIGP is substantially purified by immunoaffinity chromatography using antibodies specific for SIGP. An immunoaffinity column is constructed by covalently coupling anti-SIGP antibody to an activated chromatographic resin, such as CNBr-activated Sepharose (Pharmacia & Upjohn). After the coupling, the resin is blocked and washed according to the manufacturer's instructions.

Media containing SIGP are passed over the immunoaffinity column, and the column is washed under conditions that allow the preferential absorbance of SIGP (e.g., high ionic strength buffers in the presence of detergent). The column is eluted under conditions that disrupt antibody/SIGP binding (e.g., a buffer of pH 2 to pH 3, or a high concentration of a chaotrope, such as urea or thiocyanate ion), and SIGP is collected.

# XII. Identification of Molecules Which Interact with SIGP

SIGP, or biologically active fragments thereof, are labeled with <sup>125</sup>I Bolton-Hunter reagent. (See, e.g., Bolton et al. (1973) Biochem. J. 133:529.) Candidate molecules previously arrayed in the wells of a multi-well plate are incubated with the labeled SIGP, washed, and any wells with labeled SIGP complex are assayed. Data obtained using different concentrations of SIGP are used to calculate values for the number, affinity, and association of SIGP with the candidate molecules.

Various modifications and variations of the described methods and systems of the invention will be apparent to those skilled in the art without departing from the scope and spirit of the invention. Although the invention has been described in connection with specific preferred embodiments, it should be understood that the invention as claimed should not be

unduly limited to such specific embodiments. Indeed, various modifications of the described modes for carrying out the invention which are obvious to those skilled in molecular biology or related fields are intended to be within the scope of the following claims.

#### SEQUENCE LISTING

- (1) GENERAL INFORMATION:
  - (i) APPLICANT: Lal, Preeti
    Hillman, Jennifer L.
    Corley, Neil C.
    Guegler, Karl J.
    Baugh, Mariah
    Sather, Susan
    Shah, Purvi
  - (ii) TITLE OF INVENTION: HUMAN SIGNAL PEPTIDE-CONTAINING PROTEINS
  - (iii) NUMBER OF SEQUENCES:154
  - (iv) CORRESPONDENCE ADDRESS:
    - (A) ADDRESSEE: INCYTE PHARMACEUTICALS, INC.
    - (B) STREET: 3174 PORTER DRIVE
    - (C) CITY: PALO ALTO
    - (D) STATE: CALIFORNIA
    - (E) COUNTRY: USA
    - (F) ZIP: 94304
  - (v) COMPUTER READABLE FORM:
    - (A) MEDIUM TYPE: Floppy disk
    - (B) COMPUTER: IBM PC compatible
    - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
    - (D) SOFTWARE: Word Perfect 6.1 for Windows/MS-DOS 6.2
  - (vi) CURRENT APPLICATION DATA:
    - (A) APPLICATION NUMBER: TO BE ASSIGNED
    - (B) FILING DATE: HEREWITH
    - (C) CLASSIFICATION:
  - (viii) ATTORNEY/AGENT INFORMATION:
    - (A) NAME: BILLINGS, LUCY J.
    - (B) REGISTRATION NUMBER: 36,749
    - (C) REFERENCE/DOCKET NUMBER: PF-0459 US
    - (ix) TELECOMMUNICATION INFORMATION:
      - (A) TELEPHONE: (650) 855-0555
      - (B) TELEFAX: (650) 845-4166
- (2) INFORMATION FOR SEQ ID NO: 1:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 348 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:

#### PF-0459 US

(A) LIBRARY: HEARNOT01

(B) CLONE: 305841

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

```
Met Ala Ala Thr Leu Gly Pro Leu Gly Ser Trp Gln Gln Trp Arg
                                     10
Arg Cys Leu Ser Ala Arg Asp Gly Ser Arg Met Leu Leu Leu
                 20
                                     25
Leu Leu Gly Ser Gly Gln Gly Pro Gln Gln Val Gly Ala Gly
                                     40
                 35
Gln Thr Phe Glu Tyr Leu Lys Arg Glu His Ser Leu Ser Lys Pro
                 50
                                     55
Tyr Gln Gly Val Gly Thr Gly Ser Ser Ser Leu Trp Asn Leu Met
                 65
                                     70
Gly Asn Ala Met Val Met Thr Gln Tyr Ile Arg Leu Thr Pro Asp
                 80
                                     85
Met Gln Ser Lys Gln Gly Ala Leu Trp Asn Arg Val Pro Cys Phe
                                    100
                 95
Leu Arg Asp Trp Glu Leu Gln Val His Phe Lys Ile His Gly Gln
                110
                                    115
                                                        120
Gly Lys Lys Asn Leu His Gly Asp Gly Leu Ala Ile Trp Tyr Thr
                125
                                    130
Lys Asp Arg Met Gln Pro Gly Pro Val Phe Gly Asn Met Asp Lys
                140
                                    145
Phe Val Gly Leu Gly Val Phe Val Asp Thr Tyr Pro Asn Glu Glu
                155
                                    160
Lys Gln Gln Glu Arg Val Phe Pro Tyr Ile Ser Ala Met Val Asn
                170
                                    175
Asn Gly Ser Leu Ser Tyr Asp His Glu Arg Asp Gly Arg Pro Thr
                                    190
                185
Glu Leu Gly Gly Cys Thr Ala Ile Val Arg Asn Leu His Tyr Asp
                200
                                    205
                                                         210
Thr Phe Leu Val Ile Arg Tyr Val Lys Arg His Leu Thr Ile Met
                215
                                    220
                                                        225
Met Asp Ile Asp Gly Lys His Glu Trp Arg Asp Cys Ile Glu Val
                230
                                    235
Pro Gly Val Arg Leu Pro Arg Gly Tyr Tyr Phe Gly Thr Ser Ser
                                    250
                                                         255
Ile Thr Gly Asp Leu Ser Asp Asn His Asp Val Ile Ser Leu Lys
                                    265
                260
Leu Phe Glu Leu Thr Val Glu Arg Thr Pro Glu Glu Glu Lys Leu
                                    280
                275
His Arg Asp Val Phe Leu Pro Ser Val Asp Asn Met Lys Leu Pro
                                    295
                290
                                                         300
Glu Met Thr Ala Pro Leu Pro Pro Leu Ser Gly Leu Ala Leu Phe
                305
                                    310
                                                         315
Leu Ile Val Phe Phe Ser Leu Val Phe Ser Val Phe Ala Ile Val
                                    325
                320
Ile Gly Ile Ile Leu Tyr Asn Lys Trp Gln Glu Gln Ser Arg Lys
                                    340
                                                         345
Arg Phe Tyr
```

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 194 amino acids
  - (B) TYPE: amino acid

- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: EOSIHET02
- (B) CLONE: 322866

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:

```
Met Gly Met Ser Ser Leu Lys Leu Leu Lys Tyr Val Leu Phe Phe
                                     10
Phe Asn Leu Leu Phe Trp Ile Cys Gly Cys Cys Ile Leu Gly Phe
                                     25
                 20
Gly Ile Tyr Leu Leu Ile His Asn Asn Phe Gly Val Leu Phe His
                 35
                                     40
Asn Leu Pro Ser Leu Thr Leu Gly Asn Val Phe Val Ile Val Gly
                                     55
                                                          60
                 50
Ser Ile Ile Met Val Val Ala Phe Leu Gly Cys Met Gly Ser Ile
                                     70
                 65
Lys Glu Asn Lys Cys Leu Leu Met Ser Phe Phe Ile Leu Leu
                                     85
                 80
Ile Ile Leu Leu Ala Glu Val Thr Leu Ala Ile Leu Leu Phe Val
                                    100
                 95
Tyr Glu Gln Lys Leu Asn Glu Tyr Val Ala Lys Gly Leu Thr Asp
                                                         120
                                    115
                110
Ser Ile His Arg Tyr His Ser Asp Asn Ser Thr Lys Ala Ala Trp
                                                         135
                                    130
                125
Asp Ser Ile Gln Ser Phe Leu Gln Cys Cys Gly Ile Asn Gly Thr
                                    145
                140
Ser Asp Leu Asp Ser Gly Ser Pro Ala Ser Cys Pro Ser Asp Arg
                                    160
                155
Lys Val Glu Gly Cys Tyr Ala Lys Glu Asp Phe Gly Phe Ile Gln
                170
                                    175
Phe Pro Val Tyr Arg Asn His His His Leu Cys Met Cys Asp
                                    190
                185
```

- (2) INFORMATION FOR SEQ ID NO:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 342 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: BEPINOT01
    - (B) CLONE: 546656

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:

```
Cys Arg Asn Thr Phe Asp His Pro Tyr Pro Thr Thr Lys Leu Met
                 65
                                     70
Trp Ile Pro Asp Thr Lys Gly Val Tyr Pro Asp Leu Leu Ala Thr
                                     85
                 80
Ser Gly Asp Tyr Leu Arg Val Trp Arg Val Gly Glu Thr Glu Thr
                 95
                                    100
                                                         105
Arg Leu Glu Cys Leu Leu Asn Asn Lys Asn Ser Asp Phe Cys
                110
                                    115
Ala Pro Leu Thr Ser Phe Asp Trp Asn Glu Val Asp Pro Tyr Leu
                                    130
                125
Leu Gly Thr Ser Ser Ile Asp Thr Thr Cys Thr Ile Trp Gly Leu
                140
                                    145
Glu Thr Gly Gln Val Leu Gly Arg Val Asn Leu Val Ser Gly His
                155
                                    160
Val Lys Thr Gln Leu Ile Ala His Asp Lys Glu Val Tyr Asp Ile
                170
                                    175
Ala Phe Ser Arg Ala Gly Gly Gly Arg Asp Met Phe Ala Ser Val
                185
                                    190
                                                         195
Gly Ala Asp Gly Ser Val Arg Met Phe Asp Leu Arg His Leu Glu
                200
                                    205
                                                         210
His Ser Thr Ile Ile Tyr Glu Asp Pro Gln His His Pro Leu Leu
                                    220
                215
Arg Leu Cys Trp Asn Lys Gln Asp Pro Asn Tyr Leu Ala Thr Met
                230
                                    235
Ala Met Asp Gly Met Glu Val Val Ile Leu Asp Val Arg Val Pro
                245
                                    250
Cys Thr Pro Val Ala Arg Leu Asn Asn His Arg Ala Cys Val Asn
                260
                                    265
Gly Ile Ala Trp Ala Pro His Ser Ser Cys His Ile Cys Thr Ala
                275
                                    280
Ala Asp Asp His Gln Ala Leu Ile Trp Asp Ile Gln Gln Met Pro
                290
                                    295
Arg Ala Ile Glu Asp Pro Ile Leu Ala Tyr Thr Ala Glu Gly Glu
                                    310
                305
Ile Asn Asn Val Gln Trp Ala Ser Thr Gln Pro Asp Trp Ile Ala
                320
                                    325
Ile Cys Tyr Asn Asn Cys Leu Glu Ile Leu Arg Val
                335
```

#### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 656 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: SYNORAT03
- (B) CLONE: 693453

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

				35					40					45
				Ser 50					55				Lys	60
Val	Asp	Val	Tyr	Ser 65	Gln	Ile	Leu	Arg	Lys 70	Leu	Phe	Asn	Glu	Ser 75
His	Gly	Ile	Phe	Leu 80	Gly	Leu	Gln	Arg	Ile 85	Asp	Glu	Glu	Leu	Thr 90
Gly	Lys	Ser	Arg		Ser	Gln	Leu	Val	Arg 100	Val	Ser	Lys	Asn	Tyr 105
Arg	Ser	Val	Ile		Ala	Суз	Met	Glu	Glu 115	Met	His	Gln	Val	Ala 120
Ile	Ala	Ala	Lys	Asp 125	Pro	Ala	Asn	Gly	Arg 130	Gln	Phe	Ser	Ser	Gln 135
Val	Ser	Ile	Leu	Ser 140	Ala	Met	Glu	Leu	Ile 145	Trp	Asn	Leu	Cys	Glu 150
Ile	Leu	Phe	Ile	Glu 155	Val	Ala	Pro	Ala	Gly 160	Pro	Leu	Leu	Leu	His 165
Leu	Leu	Asp	Trp	Val 170	Arg	Leu	His	Val	Cys 175	Glu	Val	Asp	Ser	Leu 180
Ser	Ala	Asp	Val	Leu 185	Gly	Ser	Glu	Asn	Pro 190	Ser	Lys	His	Asp	Ser 195
Phe	Trp	Asn	Leu	Val 200	Thr	Ile	Leu	Val	Leu 205	Gln	Gly	Arg	Leu	Asp 210
Glu	Ala	Arg	Gln	Met 215	Leu	Ser	Lys	Glu	Ala 220	Asp	Ala	Ser	Pro	Ala 225
Ser	Ala	Gly	Ile	Cys 230	Arg	Ile	Met	Gly	Asp 235	Leu	Met	Arg	Thr	Met 240
Pro	Ile	Leu	Ser	Pro 245	Gly	Asn	Thr	Gln	Thr 250	Leu	Thr	Glu	Leu	Glu 255
Leu	Lys	Trp	Gln	His 260	Trp	His	Glu	Glu	Cys 265	Glu	Arg	Tyr	Leu	Gln 270
Asp	Ser	Thr	Phe		Thr	Ser	Pro	His	Leu 280	Glu	Ser	Leu	Leu	Lys 285
				Asp 290					295				Glu	300
				305					310				Tyr	315
				320					325				Gln	330
				335					340					Leu 345
_				350					355					Val 360
	-			365					370					Ala 375
				380	)				385					His 390
				395					400	1				Glu 405
				410	)				415	)				Gly 420
				425	5				430	)				Leu 435
				440	)				445	;				Ala 450
				a Arg 455	g Il∈ 5				460	)				Gln 465
				470	)				475	5				y Asn 480
Ası	n Arç	y Lei	ı Gly	/ Sei 485		a Lei	ı Ser	Trp	Ser 490		e Aro	g Ala	a Lys	495

```
Ala Ala Phe Ala Thr Leu Val Ser Asp Arg Phe Leu Arg Asp Tyr
                                   505
               500
Cys Glu Arg Gly Cys Phe Ser Asp Leu Asp Leu Ile Asp Asn Leu
                                   520
               515
Gly Pro Ala Met Met Leu Ser Asp Arg Leu Thr Phe Leu Gly Lys
                                                       540
                                   535
               530
Tyr Arg Glu Phe His Arg Met Tyr Gly Glu Lys Arg Phe Ala Asp
                                   550
                                                       555
               545
Ala Ala Ser Leu Leu Ser Leu Met Thr Ser Arg Ile Ala Pro
                                                       570
                560
                                   565
Arg Ser Phe Trp Met Thr Leu Leu Thr Asp Ala Leu Pro Leu Leu
                                   580
                575
Glu Gln Lys Gln Val Ile Phe Ser Ala Glu Gln Thr Tyr Glu Leu
                                   595
                                                       600
                590
Met Arg Cys Leu Glu Asp Leu Thr Ser Arg Arg Pro Val His Gly
                                   610
                605
Glu Ser Asp Thr Glu Gln Leu Gln Asp Asp Asp Ile Glu Thr Thr
                                                       630
                                    625
                620
Lys Val Glu Met Leu Arg Leu Ser Leu Ala Arg Asn Leu Ala Arg
                                   640
                635
Ala Ile Ile Arg Glu Gly Ser Leu Glu Gly Ser
                650
```

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 236 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRAITUT03
- (B) CLONE: 866885

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

Met	Ala	Pro	Asp	Pro 5	Trp	Phe	Ser	Thr	Tyr 10	Asp	Ser	Thr	Cys	Gln 15
Ile	Ala	Gln	Glu	Ile 20	Ala	Glu	Lys	Ile	Gln 25	Gln	Arg	Asn	Gln	Tyr 30
			Gly	35					40					45
Ala	Leu	Leu	Gln	Asn 50	Leu	Lys	Glu	Lys	Ile 55	Ala	Leu	Leu	Lys	Asp 60
Leu	Leu	Leu	Arg	Ala 65	Val	Ser	Thr	His	Gln 70	Ile	Thr	Gln	Leu	Glu 75
Gly	Asp	Arg	Arg		Asn	Leu	Leu	Asp	Asp 85	Leu	Val	Thr	Arg	Glu 90
Arg	Leu	Leu	Leu		Ser	Phe	Lys	Asn	Glu 100	Gly	Ala	Glu	Pro	Asp 105
Leu	Ile	Arg	Ser	Ser 110	Leu	Met	Ser	Glu	Glu 115	Ala	Lys	Arg	Gly	Ala 120
Pro	Asn	Pro	Trp	Leu 125	Phe	Glu	Glu	Pro	Glu 130	Glu	Thr	Arg	Gly	Leu 135
Gly	Phe	Asp	Glu		Arg	Gln	Gln	Gln	Gln 145	Lys	Ile	Ile	Gln	Glu 150
Gln	Asp	Ala	Gly		Asp	Ala	Leu	Ser	Ser	Ile	Ile	Ser	Arg	Gln

				155					160					165
Lys	Gln	Met	Gly		Glu	Ile	Gly	Asn	Glu 175	Leu	Asp	Glu	Gln	Asn 180
~ 3	<b>-</b> 3	<b>-</b> 1 -	7	170	T 011	71.7.	7) cn	Tou	<b>-</b> ,	Gla	Asn	Thr	Asp	
				185					190					T 3 2
Lys	Leu	Arg	Asn		Thr	Arg	Arg	Val	Asn 205	Met	Val	Asp	Arg	Lys 210
				200		* 7	3.7 - ±	77 - 7		т о	T 011	T 011	T 011	
Ser	Ala	Ser	Cys		Met	TTE	мет	vaı	220	ьеи	Leu	пеп	пеа	225
				215			_	ъ.		70				223
Ala	Ile	Val	Val		Ala	Val	Trp	Pro	Thr	Asn				
				230					235					

# (2) INFORMATION FOR SEQ ID NO: 6:

- (i) SEQUENCE CHARACTERISTICS:

  (A) LENGTH: 195 amino acids
  (B) TYPE: amino acid
  (C) STRANDEDNESS: single
  (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGNOT03 (B) CLONE: 1242271

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:

Met	Leu	Leu	Asp	Thr 5	Val	Gln	Lys	Val	Phe 10	Gln	Lys	Met	Leu	Glu 15
Cys	Ile	Ala	Arg	Ser 20	Phe	Arg	Lys	Gln		Glu	Glu	Gly	Leu	Arg 30
Leu	Leu	Tyr	Ser	Val	Gln	Arg	Pro	Leu		Glu	Phe	Ile	Thr	Ala 45
Val	Gln	Ser	Arg		Thr	Asp	Thr	Pro	Val 55	His	Arg	Gly	Val	Leu 60
				Ala 65					70	Ile				75
Arg	Lys	Val	Ser	Asp 80	Val	Glu	Glu	Leu	Thr 85	Pro	Pro	Glu	His	Leu 90
Ser	Asp	Leu	Pro		Phe	Ser	Arg	Cys	Leu 100	Ile	Gly	Ile	Ile	Ile 105
Lys	Ser	Ser	Asn	Val	Val	Arg	Ser	Phe	Leu 115	Asp	Glu	Leu	Lys	Ala 120
Cys	Val	Ala	Ser	Asn 125	Asp	Ile	Glu	Gly	Ile 130	Val	Cys	Leu	Thr	Ala 135
Ala	Val	His	Ile	Ile 140	Leu	Val	Ile	Asn	Ala 145	Gly	Lys	His	Lys	Ser 150
Ser	Lys	Val	Arg	Glu 155	Val	Ala	Ala	Thr		His	Arg	Lys	Leu	Lys 165
Thr	Phe	Met	Glu	Ile 170	Thr	Leu	Glu	Glu		Ser	Ile	Glu	Arg	Phe 180
Leu	Tyr	Glu	Ser			Arg	Thr	Leu		Glu	Leu	Leu	Asn	Ser 195

- (2) INFORMATION FOR SEQ ID NO: 7:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 608 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single (D) TOPOLOGY: linear

- (vii) IMMEDIATE SOURCE:
   (A) LIBRARY: LUNGFET03
   (B) CLONE: 1255027
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:

Met	Thr	Lys	Thr	Asp	Glu	Thr	Thr	Leu	Val	Ala	Ser	Trp	Glu	Thr 15
_				20				Leu	Phe 25					30
				35				His	40					45
Ser	Ala	Arg	Ser	Lys 50	Met	Ala	Glu	Leu	Asn 55	Thr	His	Val	Asn	Val 60
				65				Ser	70					75
				80				Gln	85					90
=				95				Asp	100					105
	_			110				Lys	115					120
_				125				Gln	130					135
_				140				Gly	145					150
				155				Asn -	160					165
				170				Lys	175					180
				185				Glu -	190					195
				200				Leu	205					210
_				215				Gly	220					225
_				230				Pro	235					240
_				245				Ser	250					255
				260				Ala	265					270
				275	•			Gln	280					285
				290	}			Glu	295					300
				305	· )				310					Ile 315
				320	)				325					Glu 330
Glu	Ala	. Met	: Glu	335	e Leu	ı Thr	ALA	Arg	340	ьys	гĀS	ALC	. GIU	Glu 345

```
Leu Lys Arg Leu Thr Asp Leu Ala Ser Gln Met Ala Glu Met Gln
                                     355
                350
Leu Ala Glu Leu Arg Ala Glu Ile Lys His Phe Val Ser Glu Arg
                                     370
                                                         375
                365
Lys Tyr Asp Glu Glu Leu Gly Lys Ala Ala Arg Phe Ser Cys Asp
                                                         390
                                     385
                380
Ile Glu Gln Leu Lys Ala Gln Ile Met Leu Cys Gly Glu Ile Thr
                                     400
                395
His Pro Lys Asn Asn Tyr Ser Ser Arg Thr Pro Cys Ser Ser Leu
                                     415
                410
Leu Pro Leu Leu Asn Ala His Ala Ala Thr Ser Gly Lys Gln Ser
                                     430
                425
Asn Phe Ser Arg Lys Ser Ser Thr His Asn Lys Pro Ser Glu Gly
                                     445
                                                          450
                440
Lys Ala Ala Asn Pro Lys Met Val Ser Ser Leu Pro Ser Thr Ala
                                     460
                455
Asp Pro Ser His Gln Thr Met Pro Ala Asn Lys Gln Asn Gly Ser
                                                          480
                                     475
                470
Ser Asn Gln Arg Arg Phe Asn Pro Gln Tyr His Asn Asn Arg
                                     490
                485
Leu Asn Gly Pro Ala Lys Ser Gln Gly Ser Gly Asn Glu Ala Glu
                                     505
                500
Pro Leu Gly Lys Gly Asn Ser Arg His Glu His Arg Arg Gln Pro
                                                          525
                 515
                                     520
His Asn Gly Phe Arg Pro Lys Asn Lys Gly Gly Ala Lys Asn Gln
                                     535
                 530
Glu Ala Ser Leu Gly Met Lys Thr Pro Glu Ala Pro Ala His Ser
                                     550
                 545
Glu Lys Pro Arg Arg Arg Gln His Ala Ala Asp Thr Ser Glu Ala
                                     565
                 560
Arg Pro Phe Arg Gly Ser Val Gly Arg Val Ser Gln Cys Asn Leu
                                                          585
                 575
                                     580
Cys Pro Thr Arg Ile Glu Val Ser Thr Asp Ala Ala Val Leu Ser
                                     595
                                                          600
                 590
Val Pro Ala Val Thr Leu Val Ala
```

- (2) INFORMATION FOR SEQ ID NO:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 267 amino acids(B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: TESTTUT02
    - (B) CLONE: 1273453
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:

Met Val Ile Ser Trp His Leu Ala Ser Asp Met Asp Cys Val Val 5 10 15

Thr Leu Thr Thr Asp Ala Ala Arg Arg Ile Tyr Asp Glu Thr Gln 20 25 30

Gly Arg Gln Gln Val Leu Pro Leu Asp Ser Ile Tyr Lys Lys Thr 35 40 45

Leu Pro Asp Trp Lys Arg Ser Leu Pro His Phe Arg Asn Gly Lys

د بيد د د د

#### PF-0459 US

Leu Tyr Phe Lys Pro Ile Gly Asp Pro Val Phe Ala Arg Asp Leu 65 70 Leu Thr Phe Pro Asp Asn Val Glu His Cys Glu Thr Val Phe Gly 85 80 Met Leu Leu Gly Asp Thr Ile Ile Leu Asp Asn Leu Asp Ala Ala 95 100 Asn His Tyr Arg Lys Glu Val Val Lys Ile Thr His Cys Pro Thr 110 115 120 Leu Leu Thr Arg Asp Gly Asp Arg Ile Arg Ser Asn Gly Lys Phe 125 130 Gly Gly Leu Gln Asn Lys Ala Pro Pro Met Asp Lys Leu Arg Gly 145 150 140 Met Val Phe Gly Ala Pro Val Pro Lys Gln Cys Leu Ile Leu Gly 155 160 Glu Gln Ile Asp Leu Leu Gln Gln Tyr Arg Ser Ala Val Cys Lys 170 175 Leu Asp Ser Val Asn Lys Asp Leu Asn Ser Gln Leu Glu Tyr Leu 190 185 Arg Thr Pro Asp Met Arg Lys Lys Gln Glu Leu Asp Glu His 200 205 210 Glu Lys Asn Leu Lys Leu Ile Glu Glu Lys Leu Gly Met Thr Pro 215 220 225 Ile Arg Lys Cys Asn Asp Ser Leu Arg His Ser Pro Lys Val Glu 230 235 Thr Thr Asp Cys Pro Val Pro Pro Lys Arg Met Arg Arg Glu Ala 250 245 Thr Arg Gln Asn Arg Ile Ile Thr Lys Thr Asp Val 260

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 285 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: TESTTUT02
- (B) CLONE: 1275261

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9 :

Met Val Met Arg Pro Leu Trp Ser Leu Leu Trp Glu Ala Leu Leu Pro Ile Thr Val Thr Gly Ala Gln Val Leu Ser Lys Val Gly 20 2.5 Gly Ser Val Leu Leu Val Ala Ala Arg Pro Pro Gly Phe Gln Val 35 40 Arg Glu Ala Ile Trp Arg Ser Leu Trp Pro Ser Glu Glu Leu Leu 55 Ala Thr Phe Phe Arg Gly Ser Leu Glu Thr Leu Tyr His Ser Arg 65 70 Phe Leu Gly Arg Ala Gln Leu His Ser Asn Leu Ser Leu Glu Leu 80 85 Gly Pro Leu Glu Ser Gly Asp Ser Gly Asn Phe Ser Val Leu Met 100

```
Val Asp Thr Arg Gly Gln Pro Trp Thr Gln Thr Leu Gln Leu Lys
                                    115
                110
Val Tyr Asp Ala Val Pro Arg Pro Val Val Gln Val Phe Ile Ala
                                    130
                                                         135
                125
Val Glu Arg Asp Ala Gln Pro Ser Lys Thr Cys Gln Val Phe Leu
                                                         150
                                    145
                140
Ser Cys Trp Ala Pro Asn Ile Ser Glu Ile Thr Tyr Ser Trp Arg
                                                         165
                                    160
                155
Arg Glu Thr Thr Met Asp Phe Gly Met Glu Pro His Ser Leu Phe
                                                         180
                                    175
                170
Thr Asp Gly Gln Val Leu Ser Ile Ser Leu Gly Pro Gly Asp Arg
                                    190
                185
Asp Val Ala Tyr Ser Cys Ile Val Ser Asn Pro Val Ser Trp Asp
                                    205
                                                         210
                200
Leu Ala Thr Val Thr Pro Trp Asp Ser Cys His His Glu Ala Ala
                                    220
                215
Pro Gly Lys Ala Ser Tyr Lys Asp Val Leu Leu Val Val Pro
                                                         240
                                     235
                230
Val Ser Leu Leu Leu Met Leu Val Thr Leu Phe Ser Ala Trp His
                                     250
                245
Trp Cys Pro Cys Ser Gly Lys Lys Lys Asp Val His Ala Asp
                                                         270
                                     265
                260
Arg Val Gly Pro Glu Thr Glu Asn Pro Leu Val Gln Asp Leu Pro
                                     280
                275
```

- (2) INFORMATION FOR SEQ ID NO:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 76 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: COLNNOT16
    - (B) CLONE: 1281682
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:

 Met
 Pro
 Phe
 Thr
 Arg
 Pro
 Leu
 Lys
 His
 Phe
 Val
 Ser
 Leu
 His
 15

 Pro
 Ser
 Ala
 Ser
 Gln
 Val
 His
 Asn
 Ala
 Gly
 Gln
 His
 Gln
 Lys
 Leu

 Lys
 Thr
 Leu
 Gly
 Leu
 Ala
 Leu
 Gly
 Glu
 Gly
 Gly
 Arg

 Glu
 Gln
 Asn
 Leu
 Cys
 Thr
 Ser
 Leu
 Phe
 Asn
 Leu
 Glu
 Ile
 Arg
 His

 Fro
 Arg
 Asp
 Ala
 Ile
 Ile
 Phe
 Cys
 Val
 Ser
 Ile
 Val
 Val
 Pro
 Leu

 Fro
 Arg
 Asp
 Ala
 Ile
 Ile
 Phe
 Cys
 Val
 Ser
 Ile
 Val
 Pro
 Leu

 Fro
 Arg
 Arg
 Arg
 Arg
 <td

- (2) INFORMATION FOR SEQ ID NO:
  - (i) SEQUENCE CHARACTERISTICS:

128

- (A) LENGTH: 147 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRSTNOT07
- (B) CLONE: 1298305

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:

Met Thr Ala Ser Thr Gly His Leu Gly Leu Gly Trp Ser Ala Arg Pro Cys Pro Cys Gly Thr Leu Gly Ser Cys Phe Leu Ser Leu Phe 20 25 Ala Ala Leu Leu Trp Leu Ala Ala Ala Val Leu Gln Ala Cys Val 35 40 Gly His Ser Asp Glu Gly Cys Gly Ala Ser Gln Cys Arg Arg Ala 50 55 Ala Leu Gly Ile Val Pro Ser Pro Val Ser Val Leu Arg Thr Tyr 65 70 Pro Gly Leu His His Gln Asp Pro Val Phe Gly Phe Arg Arg Pro 80 85 Ser Met Gly Lys Thr Arg His Gln Pro Leu Gln Gln Trp Val Pro 95 100 Leu Ala Cys Gly His Gln Leu Gly Asp Pro Gly Ser Gly Pro Leu 110 115 Leu Ser Pro Val Ser Leu Cys Cys Gly Phe Trp Ala Val Met Ser 125 130 Pro Pro Leu Lys Asp Val Phe Thr Leu Thr Ser Gly 140

- (2) INFORMATION FOR SEQ ID NO:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 261 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: LUNGNOT12
    - (B) CLONE: 1360501

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:

Met Glu Leu Gln Val Thr Ile Leu Phe Leu Pro Ser Ile 10 Cys Ser Ser Asn Ser Thr Gly Val Leu Glu Ala Ala Asn Asn Ser 25 Leu Val Val Thr Thr Thr Lys Pro Ser Ile Thr Thr Pro Asn Thr 35 40 Glu Ser Leu Gln Lys Asn Val Val Thr Pro Thr Thr Gly Thr Thr 50 55 Pro Lys Gly Thr Ile Thr Asn Glu Leu Leu Lys Met Ser Leu Met 65 70 Ser Thr Ala Thr Phe Leu Thr Ser Lys Asp Glu Gly Leu Lys Ala 80 85

```
Thr Thr Asp Val Arg Lys Asn Asp Ser Ile Ile Ser Asn Val
                95
                                   100
Thr Val Thr Ser Val Thr Leu Pro Asn Ala Val Ser Thr Leu Gln
                                    115
                110
Ser Ser Lys Pro Lys Thr Glu Thr Gln Ser Ser Ile Lys Thr Thr
                125
                                    130
                                                        135
Glu Ile Pro Gly Ser Val Leu Gln Pro Asp Ala Ser Pro Ser Lys
                140
                                    145
Thr Gly Thr Leu Thr Ser Ile Pro Val Thr Ile Pro Glu Asn Thr
                                    160
                155
Ser Gln Ser Gln Val Ile Gly Thr Glu Gly Gly Lys Asn Ala Ser
                170
                                    175
Thr Ser Ala Thr Ser Arg Ser Tyr Ser Ser Ile Ile Leu Pro Val
                                    190
                185
Val Ile Ala Leu Ile Val Ile Thr Leu Ser Val Phe Val Leu Val
                200
                                    205
Gly Leu Tyr Arg Met Cys Trp Lys Ala Asp Pro Gly Thr Pro Glu
                                    220
                215
Asn Gly Asn Asp Gln Pro Gln Ser Asp Lys Glu Ser Val Lys Leu
                230
                                    235
Leu Thr Val Lys Thr Ile Ser His Glu Ser Gly Glu His Ser Ala
                245
                                   250
Gln Gly Lys Thr Lys Asn
                260
```

#### (2) INFORMATION FOR SEQ ID NO: 13:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 213 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGNOT12
- (B) CLONE: 1362406

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:

Met Ala Gly Cys Pro Ala Asp Arg Ser Ile Leu Ala Pro Leu Ala 10 Trp Asp Leu Gly Leu Leu Leu Phe Val Gly Gln His Ser Leu 25 20 Met Ala Ala Glu Arg Val Lys Ala Trp Thr Ser Arg Tyr Phe Gly 35 40 Val Leu Gln Arg Ser Leu Tyr Val Ala Cys Thr Ala Leu Ala Leu 50 55 Gln Leu Val Met Arg Tyr Trp Glu Pro Ile Pro Lys Gly Pro Val 65 70 Leu Trp Glu Ala Arg Ala Glu Pro Trp Ala Thr Trp Val Pro Leu 80 85 Leu Cys Phe Val Leu His Val Ile Ser Trp Leu Leu Ile Phe Ser 95 100 Ile Leu Leu Val Phe Asp Tyr Ala Glu Leu Met Gly Leu Lys Gln 110 115 Val Tyr Tyr His Val Leu Gly Leu Gly Glu Pro Leu Ala Leu Lys 130 125 Ser Pro Arg Ala Leu Arg Leu Phe Ser His Leu Arg His Pro Val

				140					145				_	150
Cys	Val	Glu	Leu	Leu 155	Thr	Val	Leu	Trp	Val 160	Val	Pro	Thr	Leu	GLy 165
Thr	Asp	Arg	Leu	Leu 170	Leu	Ala	Phe	Leu	Leu 175	Thr	Leu	Tyr	Leu	Gly 180
Leu	Ala	His	Gly	Leu 185	Asp	Gln	Gln	Asp	Leu 190	Arg	Tyr	Leu	Arg	Ala 195
Gln	Leu	Gln	Arg	Lys 200	Leu	His	Leu	Leu	Ser 205	Arg	Pro	Gln	Asp	Gly 210
Glu	Ala	Glu												

#### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 67 amino acids

  - (B) TYPE: amino acid(C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LATRTUT02
  - (B) CLONE: 1405329
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:

Met Gln Pro Arg Pro Arg Gly Arg Pro Pro Arg Thr Arg Gly Asp 10 Glu Ala Pro Gln Trp His Leu Pro Asp Ala Ala Ala Leu Leu Pro 25 Val Arg Leu Pro Leu Ala Val Leu Val Arg Gly Thr Gln Arg Pro 35 40 Glu Arg Arg Cys Gly Arg Leu Pro Ala Gly Val Pro Gly Ala 50 Ala Arg Ser Val Ala Arg Ser 65

- (2) INFORMATION FOR SEQ ID NO: 15:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 161 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: BRAINOT12
    - (B) CLONE: 1415223
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15:

Met Leu Ala Pro Gln Arg Thr Arg Ala Pro Ser Pro Arg Ala Ala 5 10 Pro Arg Pro Thr Arg Ser Met Leu Pro Ala Ala Met Lys Gly Leu 25

Gly Leu Ala Leu Leu Ala Val Leu Leu Cys Ser Ala Pro Ala His 35 Gly Leu Trp Cys Gln Asp Cys Thr Leu Thr Thr Asn Ser Ser His 55 50 Cys Thr Pro Lys Gln Cys Gln Pro Ser Asp Thr Val Cys Ala Ser 70 65 Val Arg Ile Thr Asp Pro Ser Ser Ser Arg Lys Asp His Ser Val 85 80 Asn Lys Met Cys Ala Ser Ser Cys Asp Phe Val Lys Arg His Phe 95 100 Phe Ser Asp Tyr Leu Met Gly Phe Ile Asn Ser Gly Ile Leu Lys 115 110 Val Asp Val Asp Cys Cys Glu Lys Asp Leu Cys Asn Gly Ala Ala 125 130 Gly Ala Gly His Ser Pro Trp Ala Leu Ala Gly Gly Leu Leu 140 145 Ser Leu Gly Pro Ala Leu Leu Trp Ala Gly Pro

### (2) INFORMATION FOR SEQ ID NO: 16

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 141 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: BRAINOT12
  - (B) CLONE: 1416553
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:

Met Trp Ala Gln Arg Val Leu Thr Leu Trp Gln Gly Leu Ser Trp 10 15 Gly Arg Pro Pro Ser Gly Pro Gly Ala Met Ala Pro Arg Gly Gln 25 Ala Asp Leu Leu Pro Ala Val Ser Thr Pro Phe Leu Ile Thr Val 35 Trp Ser Pro Ser Phe Gly Cys Ser Leu Arg Cys Val Leu Gly Ser 50 55 Ser Glu Pro Glu Ala Ser Phe Trp Lys Pro Ala Val Leu Pro Ala 65 70 Pro Val Gln Lys Pro Leu Ser Pro Ala Phe Pro Gln Ala Gly Val 80 85 Gly Val Gly Gly Leu Cys Pro Ser Ser Leu Thr Leu Glu Arg Trp 95 100 105 Glu Ala Gly Asn Leu His Leu Gly Ala Trp Ala Pro Pro Leu Cys 110 115 120 Ala Ser Gly Phe Pro Ala Pro Gly Arg Gly Cys Ser Pro Ser Trp 125 130 135 Thr Pro Ala Cys Pro Ser 140

(2) INFORMATION FOR SEQ ID NO: 1

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 152 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: KIDNNOT09
  - (B) CLONE: 1418517
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:

Met Glu Asp Glu Glu Val Ala Glu Ser Trp Glu Glu Ala Ala Asp 10 Ser Gly Glu Ile Asp Arg Arg Leu Glu Lys Lys Leu Lys Ile Thr 20 25 Gln Lys Glu Ser Arg Lys Ser Lys Ser Pro Pro Lys Val Pro Ile 35 40 Val Ile Gln Asp Asp Ser Leu Pro Ala Gly Pro Pro Pro Gln Ile 50 55 Arg Ile Leu Lys Arg Pro Thr Ser Asn Gly Val Val Ser Ser Pro 65 70 Asn Ser Thr Ser Arg Pro Thr Leu Pro Val Lys Ser Leu Ala Gln 80 85 90 Arg Glu Ala Glu Tyr Ala Glu Ala Arg Lys Arg Ile Leu Gly Ser 95 100 105 Ala Ser Pro Glu Glu Glu Glu Lys Pro Ile Leu Asp Arg Pro 110 115 Thr Arg Ile Ser Gln Pro Glu Asp Ser Arg Gln Pro Asn Asn Val 125 130 Ile Arg Gln Pro Leu Gly Pro Asp Gly Ser Gln Gly Phe Lys Gln 140 145 Arg Arg

- (2) INFORMATION FOR SEQ ID NO:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 742 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: PANCNOTO8
    - (B) CLONE: 1438165
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:

_		_		65					70	_				75
				80					85				Leu	90
Asn	Ser	Val	Gln	Ile 95	Arg	Ile	Lys	Leu	Met 100	Ala	Glu	Gly	Glu	Leu 105
Lys	Asp	Met	Glu	Gln 110	Phe	Phe	Asp	Asp	Leu 115	Ser	Asp	Ser	Phe	Ser 120
Gly	Thr	Glu	Pro	Glu 125	Val	His	Lys	Thr	Ser 130	Val	Val	Gly	Leu	
Leu	Arg	His	Met		Leu	Ala	Tyr	Ser		Leu	Ser	Phe	Ser	
Val	Phe	Lys	Leu		Thr	Ala	Leu	Gln		Tyr	Phe	Gln	Asn	
Glu	Lys	Lys	Thr		Glu	Asp	Ala	Asp		Glu	Leu	Thr	Ser	
Asp	Glu	Gly	Glu		Lys	Met	Glu	Lys		Glu	Leu	Asp	Val	
Val	Arg	Glu	Glu		Val	Ser	Cys	Ser		Pro	Leu	Ser	Gln	
Gln	Ala	Glu	Phe		Leu	Ser	Gln	Gln		Ser	Leu	Leu	Lys	
Asp	Glu	Thr	Lys		Leu	Thr	Pro	Ala		Leu	Gln	Lys	Glu	
Asn	Asn	Leu	Leu		Phe	Asn	Pro	Asp		Ala	Glu	Ala	His	
Leu	Ser	Tyr	Leu		Asn	Leu	Arg	Val		Asp	Val	Phe	Ser	Ser 270
Thr	His	Ser	Leu	Leu 275	His	Tyr	Phe	Asp		Leu	Ile	Leu	Thr	
Ala	Glu	Ser	Lys		Asn	Gly	Glu	Glu		Tyr	Gly	Arg	Ser	Leu 300
Arg	Tyr	Ala	Ala		Asn	Leu	Ala	Ala		His	Cys	Arg	Phe	
His	Tyr	Gln	Gln		Glu	Leu	Ala	Leu		Glu	Ala	Ile	Arg	
Ala	Gln	Glu	Ser		Asp	His	Val	Cys		Gln	His	Cys	Leu	
Trp	Leu	Tyr	Val	Leu 350	Gly	Gln	Lys	Arg	Ser 355	Asp	Ser	Tyr	Val	
Leu	Glu	His	Ser		Lys	Lys	Ala	Val		Phe	Gly	Leu	Pro	
Ala	Phe	Ala	Gly	Lys 380	Thr	Ala	Asn	Lys		Met	Asp	Ala	Leu	
Asp	Ser	Asp	Leu	Leu 395	His	Trp	Lys	His	Ser 400	Leu	Ser	Glu	Leu	Ile 405
Asp	Ile	Ser	Ile		Gln	Lys	Thr	Ala		Trp	Arg	Leu	Tyr	
Arg	Ser	Thr	Met		Leu	Gln	Gln	Ala		Met	Leu	Leu	Ser	
Asn	Ser	Leu	Glu		Val	Asn	Ala	Gly		Gln	Gln	Asn	Asn	
Glu	Ser	Phe	Ala		Ala	Leu	Cys	His		Ala	Glu	Leu	His	
Glu	Gln	Gly	Cys		Ala	Ala	Ala	Ser		Val	Leu	Lys	His	
Lys	Glu	Arg	Phe		Pro	Asn	Ser	Gln		Ala	Gln	Leu	Trp	
Leu	Cys	Asp	Gln		Ile	Gln	Phe	Asp		Ala	Met	Asn	Asp	
Lys	Tyr	His	Leu		Asp	Ser	Leu	Val		Gly	Ile	Thr	Ala	

```
Asn Ser Ile Glu Gly Val Tyr Arg Lys Ala Val Val Leu Gln Ala
                                    535
                530
Gln Asn Gln Met Ser Glu Ala His Lys Leu Leu Gln Lys Leu Leu
                545
                                    550
Val His Cys Gln Lys Leu Lys Asn Thr Glu Met Val Ile Ser Val
                560
                                    565
                                                         570
Leu Leu Ser Val Ala Glu Leu Tyr Trp Arg Ser Ser Ser Pro Thr
                                    580
                575
Ile Ala Leu Pro Met Leu Leu Gln Ala Leu Ala Leu Ser Lys Glu
                590
                                    595
Tyr Arg Leu Gln Tyr Leu Ala Ser Glu Thr Val Leu Asn Leu Ala
                                    610
                605
Phe Ala Gln Leu Ile Leu Gly Ile Pro Glu Gln Ala Leu Ser Leu
                620
                                    625
Leu His Met Ala Ile Glu Pro Ile Leu Ala Asp Gly Ala Ile Leu
                                    640
                635
Asp Lys Gly Arg Ala Met Phe Leu Val Ala Lys Cys Gln Val Ala
                                     655
                650
Ser Ala Ala Ser Tyr Asp Gln Pro Lys Lys Ala Glu Ala Leu Glu
                                    670
                                                         675
                665
Ala Ala Ile Glu Asn Leu Asn Glu Ala Lys Asn Tyr Phe Ala Lys
                                    685
                680
                                                         690
Val Asp Cys Lys Glu Arg Ile Arg Asp Val Val Tyr Phe Gln Ala
                                                         705
                                    700
                695
Arg Leu Tyr His Thr Leu Gly Lys Thr Gln Glu Arg Asn Arg Cys
                710
                                    715
Ala Met Leu Phe Arg Gln Leu His Gln Glu Leu Pro Ser His Gly
                                    730
                                                         735
                725
Val Pro Leu Ile Asn His Leu
```

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 805 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: THYRNOT03
- (B) CLONE: 1440381

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 19:

Met Asp Gly Ile Leu Asp Glu Ser Leu Leu Glu Thr Cys Pro Ile 10 Gln Ser Pro Leu Gln Val Phe Ala Gly Met Gly Gly Leu Ala Leu 25 20 Ile Ala Glu Arg Leu Pro Met Leu Tyr Pro Glu Val Ile Gln Gln 40 45 35 Val Ser Ala Pro Val Val Thr Ser Thr Thr Gln Glu Lys Pro Tyr 60 5.5 50 Asp Ser Asp Gln Phe Glu Trp Val Thr Ile Glu Gln Ser Gly Glu 65 Leu Val Tyr Glu Ala Pro Glu Thr Val Ala Ala Glu Pro Pro Pro 8.5 80 Ile Lys Ser Ala Val Gln Thr Met Ser Pro Ile Pro Ala His Ser

19:

T	70.7	70.7 -	Dl.	95	T .	ъ.	<del>-</del>	_	100	_	~ 1	_		105
				110					115				Ala	120
				125					130				Arg	135
Val	Leu	Gly	Val	Thr 140	Asp	Asp	Gly	Glu	Gly 145	Ser	His	Ile	Leu	Gln 150
Ser	Pro	Ser	Ala	Asn 155	Val	Leu	Pro	Thr	Leu 160	Pro	Phe	His	Val	
Arg	Ser	Leu	Phe		Thr	Thr	Pro	Leu		Thr	Asp	Asp	Gly	
Leu	Leu	Arg	Arg		Ala	Leu	Glu	Ile		Ala	Leu	His	Leu	
Leu	Val	Cys	Leu		Ala	Leu	Ser	His		Ser	Pro	Arg	Val	
Asn	Ser	Ser	Val		Gln	Thr	Glu	Pro		Val	Ser	Ser	Ser	
Asn	Pro	Thr	Ser	Thr 230	Glu	Glu	Gln	Gln		Tyr	Trp	Ala	Lys	
Thr	Gly	Phe	Gly	Thr 245	Gly	Ser	Thr	Ala		Gly	Trp	Asp	Val	
Gln	Ala	Leu	Thr	Lys 260	Gln	Arg	Leu	Glu		Glu	His	Val	Thr	
Leu	Leu	Gln	Val	Leu 275	Ala	Ser	Tyr	Ile		Pro	Val	Ser	Ser	
Val	Asn	Gly	Glu	Ala 290	Gln	Ser	Ser	His		Thr	Arg	Gly	Gln	
Ser	Asn	Ala	Leu	Pro 305	Ser	Val	Leu	Leu		Leu	Leu	Ser	Gln	Ser 315
Cys	Leu	Ile	Pro	Ala 320	Met	Ser	Ser	Tyr		Arg	Asn	Asp	Ser	
Leu	Asp	Met	Ala	Arg 335	His	Val	Pro	Leu		Arg	Ala	Leu	Leu	
Leu	Leu	Arg	Ala	Ile 350	Ala	Ser	Cys	Ala	Ala 355	Met	Val	Pro	Leu	
Leu	Pro	Leu	Ser	Thr 365	Glu	Asn	Gly	Glu		Glu	Glu	Glu	Gln	
Glu	Cys	Gln	Thr	Ser 380	Val	Gly	Thr	Leu		Ala	Lys	Met	Lys	Thr 390
Cys	Val	Asp	Thr	Tyr 395	Thr	Asn	Arg	Leu		Ser	Lys	Arg	Glu	
Val	Lys	Thr	Gly	Val 410	Lys	Pro	Asp	Ala		Asp	Gln	Glu	Pro	Glu 420
Gly	Leu	Thr	Leu		Val	Pro	Asp	Ile		Lys	Thr	Ala	Glu	
Val	Tyr	Ala	Ala		Thr	Ser	Leu	Arg		Ala	Asn	Gln	Glu	Lys 450
Asn	Trp	Val	Asn		Pro	Arg	Arg	Arg		Met	Asn	Pro	Lys	Pro 465
Leu	Ser	Val	Leu		Ser	Leu	Glu	Glu		Tyr	Val	Ala	Val	
Lys	Lys	Leu	Gln		Asp	Thr	Phe	Glu		Val	Ser	Glu	Asp	Glu
Asp	Gly	Lys	Leu		Phe	Lys	Val	Asn		His	Tyr	Met	Ser	495 Gln 510
Val	Lys	Asn	Ala		Asp	Ala	Asn	Ser		Ala	Arg	Ala	Arg	Arg
Leu	Ala	Gln	Glu		Val	Thr	Leu	Ser		Ser	Leu	Pro	Leu	
Ser	Ser	Ser	Ser		Phe	Val	Arg	Cys		Glu	Glu	Arg	Leu	540 Asp 555
				J 1 J					550					JJJ

```
Ile Met Lys Val Leu Ile Thr Gly Pro Ala Asp Thr Pro Tyr Ala
                                    565
                560
Asn Gly Cys Phe Glu Phe Asp Val Tyr Phe Pro Gln Asp Tyr Pro
                                     580
                575
Ser Ser Pro Pro Leu Val Asn Leu Glu Thr Thr Gly Gly His Ser
                                     595
                590
Val Arg Phe Asn Pro Asn Leu Tyr Asn Asp Gly Lys Val Cys Leu
                                     610
                605
Ser Ile Leu Asn Thr Trp His Gly Arg Pro Glu Glu Lys Trp Asn
                                     625
                620
Pro Gln Thr Ser Ser Phe Leu Gln Val Leu Val Ser Val Gln Ser
                                     640
                635
Leu Ile Leu Val Ala Glu Pro Tyr Phe Asn Glu Pro Gly Tyr Glu
                                     655
                650
Arg Ser Arg Gly Thr Pro Ser Gly Thr Gln Ser Ser Arg Glu Tyr
                                                          675
                                     670
                665
Asp Gly Asn Ile Arg Gln Ala Thr Val Lys Trp Ala Met Leu Glu
                                     685
                680
Gln Ile Arg Asn Pro Ser Pro Cys Phe Lys Glu Val Ile His Lys
                                     700
                                                         705
                 695
His Phe Tyr Leu Lys Arg Val Glu Ile Met Ala Gln Cys Glu Glu
                                                          720
                                     715
                710
Trp Ile Ala Asp Ile Gln Gln Tyr Ser Ser Asp Lys Arg Val Gly
                                                          735
                                     730
                 725
Arg Thr Met Ser His His Ala Ala Ala Leu Lys Arg His Thr Ala
                                     745
                 740
Gln Leu Arg Glu Glu Leu Leu Lys Leu Pro Cys Pro Glu Gly Leu
                                     760
                 755
Asp Pro Asp Thr Asp Asp Ala Pro Glu Val Cys Arg Ala Thr Thr
                                     775
                 770
Gly Ala Glu Glu Thr Leu Met His Asp Gln Val Lys Pro Ser Ser
                                                          795
                                     790
                 785
Ser Lys Glu Leu Pro Ser Asp Phe Gln Leu
                 800
```

# (2) INFORMATION FOR SEQ ID NO:

20:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 195 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGNOT14
- (B) CLONE: 1510839

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:

 Met
 Lys
 Ala
 Ser
 Gln
 Cys
 Cys
 Cys
 Leu
 Ser
 His
 Leu
 Leu
 Ala

 Ser
 Val
 Leu
 Leu
 Leu
 Leu
 Leu
 Leu
 Pro
 Glu
 Leu
 Ser
 Gly
 Pro
 Leu

 Ala
 Val
 Leu
 Leu
 Leu
 Leu
 Leu
 Ala
 Ala
 Ala
 Ala
 Pro
 Gly
 Leu
 Gly
 Pro

 Ala
 Ala
 Ala
 Ala
 Ala
 Ala
 Ala
 Pro
 Pro
 Ala
 A

				65					70					75
Pro	Arg	Gly	Ser	Glu 80	Gly	Gly	Asn	Gly	Ser 85	Asn	Pro	Val	Ala	Gly 90
Leu	Glu	Thr	Asp	Asp 95	His	Gly	Gly	Lys	Ala 100	Gly	Glu	Gly	Ser	Val 105
Gly	Gly	Gly	Leu	Ala 110	Val	Ser	Pro	Asn	Pro 115	Gly	Asp	Lys	Pro	Met 120
Thr	Gln	Arg	Ala	Leu 125	Thr	Val	Leu	Met	Val 130	Val	Ser	Gly	Ala	Val 135
Leu	Val	Tyr	Phe	Val 140	Val	Arg	Thr	Val	Arg 145	Met	Arg	Arg	Arg	Asn 150
Arg	Lys	Thr	Arg	Arg 155	Tyr	Gly	Val	Leu	Asp 160	Thr	Asn	Ile	Glu	Asn 165
Met	Glu	Leu	Thr	Pro	Leu	Glu	Gln	Asp	Asp 175	Glu	Asp	Asp	Asp	Asn 180
Thr	Leu	Phe	Asp	Ala 185	Asn	His	Pro	Arg	Arg 190	Arg	Glu	Cys	Ala	Phe 195

# (2) INFORMATION FOR SEQ ID NO: 21:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 161 amino acids

  - (B) TYPE: amino acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: SPLNNOT04
- (B) CLONE: 1534876

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

Met	Trp	Phe	Leu	Gly 5	Cys	Thr	Gly	Pro	Gly 10	Cys	Gly	Cys	Ala	Gly 15
Val	Cys	Lys	Val	Val 20	Pro	Cys	Ile	Ser	Thr 25	Gly	Phe	Glu	Thr	Ser 30
Gly	Pro	Cys	Pro	Ser 35	Ser	Arg	Glu	Gly	Phe 40	Leu	Phe	Phe	Leu	Thr 45
Gln	Val	Thr	Phe	Gln 50	Pro	Phe	Gln	Phe	Pro 55	Ser	Phe	Ser	Ala	Leu 60
Pro	Ser	Asn	Ser	Ala 65	Asn	Pro	Gly	Val	Gly 70	Ser	Gln	Gly	Gly	Arg 75
Glu	Cys	Pro	Thr	Thr 80	Phe	Ser	Gly	Gln	Pro 85	Leu	Thr	Pro	Lys	Pro 90
Leu	Pro	Pro	Ser	Ile 95	Leu	His	Pro	Leu	Pro 100	Ile	Gln	Pro	Lys	Cys 105
Pro	Gln	Leu	Gly	Leu 110	Ser	Cys	Ile	Pro	Val 115	Glu	Gly	Pro	Leu	Pro 120
Cys	Leu	Ser	Glu	Val 125	Arg	Leu	Cys	Cys	Val 130	Met	Gly	Arg	Leu	Cys 135
Pro	Ser	Pro	Pro	Leu 140	Ala	Arg	Cys	Thr	Cys 145	Phe	Leu	Val	Cys	Thr 150
Arg	Cys	Pro	Gly	Gly 155	Pro	Ser	Leu	Pro	Cys 160	Gln				

- (2) INFORMATION FOR SEQ ID NO: 22
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 160 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: SPLNNOT04
    - (B) CLONE: 1559131
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:

Met Asp Lys Leu Lys Lys Val Leu Ser Gly Gln Asp Thr Glu Asp 10 Arg Ser Gly Leu Ser Glu Val Val Glu Ala Ser Ser Leu Ser Trp 20 25 30 Ser Thr Arg Ile Lys Gly Phe Ile Ala Cys Phe Ala Ile Gly Ile 40 35 Leu Cys Ser Leu Leu Gly Thr Val Leu Leu Trp Val Pro Arg Lys 60 50 Gly Leu His Leu Phe Ala Val Phe Tyr Thr Phe Gly Asn Ile Ala 70 65 Ser Ile Gly Ser Thr Ile Phe Leu Met Gly Pro Val Lys Gln Leu 80 85 Lys Arg Met Phe Glu Pro Thr Arg Leu Ile Ala Thr Ile Met Val 100 95 Leu Leu Cys Phe Ala Leu Thr Leu Cys Ser Ala Phe Trp Trp His 110 115 Asn Lys Gly Leu Ala Leu Ile Phe Cys Ile Leu Gln Ser Leu Ala 130 125 Leu Thr Trp Tyr Ser Leu Ser Phe Ile Pro Phe Ala Arg Asp Ala 140 145 150 Val Lys Lys Cys Phe Ala Val Cys Leu Ala 155

- (2) INFORMATION FOR SEQ ID NO: 23
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 76 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: BLADNOT03
    - (B) CLONE: 1601473
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:

 Met Gln Ala Lys Tyr Ser Ser Thr Arg Asp Met Leu Asp Asp Asp
 5
 10
 15

 Gly Asp Thr Thr Met Ser Leu His Ser Gln Ala Ser Ala Thr Thr 20
 25
 30

 Arg His Pro Glu Pro Arg Arg Thr Glu His Arg Ala Pro Ser Ser 35
 40
 45

 Thr Trp Arg Pro Val Ala Leu Thr Leu Leu Thr Leu Cys Leu Val 50
 55
 60

Leu Leu Ile Gly Leu Ala Ala Leu Gly Leu Leu Cys Lys Ser Ala  $65 \hspace{1cm} 70 \hspace{1cm} 75$ 

Leu

- (2) INFORMATION FOR SEQ ID NO: 24:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 336 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: BRAITUT12
    - (B) CLONE: 1615809
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:

Met	Ile	Ser	Tyr	Ile 5	Val	Leu	Leu	Ser	Ile 10	Leu	Leu	Trp	Pro	Leu 15
Val	Val	Tyr	His	Glu 20	Leu	Ile	Gln	Arg	Met 25	Tyr	Thr	Arg	Leu	Glu 30
Pro	Leu	Leu	Met	Gln 35	Leu	Asp	Tyr	Ser	Met 40	Lys	Ala	Glu	Ala	Asn 45
				Lys 50					55					60
				Gly 65					70					75
				Leu 80					85					90
_				Ala 95					100					105
				Ile 110					115					120
				Gln 125					130					135
				Ser 140					145					150
				Glu 155					160					165
_	_			Ala 170					175					180
				Val 185					190					195
				Phe 200					205					210
				Val 215					220					225
				Thr 230					235					240
				Pro 245					250					255
				Leu 260					265					270
				Glu 275					280					285
				Gln 290					295					300
_				Glu 305					310					315
Leu	Gly	Pro	Asp	Ile	His	Ser	Leu	Val	Gln	Ser	Asp	Gln	Glu	Ala

320 Gln Ala Val Ala Glu Pro 335 325

330

(2) INFORMATION FOR SEQ ID NO: 25:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 150 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: COLNNOT19
  - (B) CLONE: 1634813
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 25:

Met Asn Leu Trp Leu Leu Ala Cys Leu Val Ala Gly Phe Leu Gly 15 10 Ala Trp Ala Pro Ala Val His Ala Gln Gly Val Phe Glu Asp Cys Cys Leu Ala Tyr His Tyr Pro Ile Gly Trp Ala Val Leu Arg Arg 35 40 Ala Trp Thr Tyr Arg Ile Gln Glu Val Ser Gly Ser Cys Asn Leu 55 50 Pro Ala Ala Ile Phe Tyr Leu Pro Lys Arg His Arg Lys Val Cys 70 65 Gly Asn Pro Lys Ser Arg Glu Val Gln Arg Ala Met Lys Leu Leu 85 80 Asp Ala Arg Asn Lys Val Phe Ala Lys Leu Arg His Asn Thr Gln 100 95 Thr Phe Gln Ala Gly Pro His Ala Val Lys Lys Leu Ser Ser Gly 115 110 Asn Ser Lys Leu Ser Ser Ser Lys Phe Ser Asn Pro Ile Ser Ser 130 125 Ser Lys Arg Asn Val Ser Leu Leu Ile Ser Ala Asn Ser Gly Leu 145

- (2) INFORMATION FOR SEQ ID NO: 26:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 217 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: UTRSNOT06
    - (B) CLONE: 1638407
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26:

Met Ala Pro Pro Ala Leu Gln Arg Gly Gln Arg Val Ala Ala Val 5 10 15

Ala Val Gly Ser Gln Ala Val Leu Gln Ile Leu Ser Arg Val Ser 25 20 Gly Arg Gln Ala Pro Pro Gln Pro Ser Gly Ser Gly Val Gly 35 Ala Gly Pro Val Val Pro Asp Gly Gly Glu Gly Pro Gln 50 55 Pro His Pro Ser Ser Ser Gln Ser Pro Pro Asp Leu Pro Leu Lys 65 Ala Gly Asp Thr Val Met Gly Lys Gln Ala Gln Arg Asp Ile Arg 80 85 Leu Arg Val Arg Ala Glu Tyr Cys Glu His Gly Pro Ala Leu Glu 95 100 Gln Gly Val Ala Ser Arg Arg Pro Gln Ala Leu Ala Arg Gln Leu 110 115 Asp Val Phe Gly Gln Ala Thr Ala Val Leu Arg Ser Arg Asp Leu 130 125 135 Gly Ser Val Val Cys Asp Ile Lys Phe Ser Glu Leu Ser Tyr Leu 140 145 Asp Ala Phe Trp Gly Asp Tyr Leu Ser Gly Ala Leu Leu Gln Ala 155 160 Leu Arg Gly Val Phe Leu Thr Glu Ala Leu Arg Glu Ala Val Gly 170 175 Arg Glu Ala Val Arg Leu Leu Val Ser Val Asp Glu Ala Asp Tyr 190 195 185 Glu Ala Gly Arg Arg Leu Leu Leu Met Ala Glu Glu Gly Gly 200 205 210 Arg Arg Pro Thr Glu Ala Ser 215

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 504 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: PROSTUT08
  - (B) CLONE: 1653112

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:

Met Ser Gln Pro Arg Thr Pro Glu Gln Ala Leu Asp Thr Pro Gly 10 Asp Cys Pro Pro Gly Arg Asp Glu Asp Ala Gly Glu Gly Ile 20 Gln Cys Ser Gln Arg Met Leu Ser Phe Ser Asp Ala Leu Leu Ser 40 35 Ile Ile Ala Thr Val Met Ile Leu Pro Val Thr His Thr Glu Ile 50 55 Ser Pro Glu Gln Gln Phe Asp Arg Ser Val Gln Arg Leu Leu Ala 70 65 Thr Arg Ile Ala Val Tyr Leu Met Thr Phe Leu Ile Val Thr Val 80 85 Ala Trp Ala Ala His Thr Arg Leu Phe Gln Val Val Gly Lys Thr 95 100 Asp Asp Thr Leu Ala Leu Leu Asn Leu Ala Cys Met Met Thr Ile

				110					115				_	120
Thr	Phe	Leu	Pro	Tyr 125	Thr	Phe	Ser	Leu	Met 130	Val	Thr	Phe	Pro	Asp 135
			Gly	Ile 140					Val 145					120
Gly	Val	Val	Gln	Ala 155	Leu	Ile	Val	Gly	Tyr 160	Ala	Phe	His	Phe	Pro 165
His	Leu	Leu	Ser	Pro 170	Gln	Ile	Gln	Arg	Ser 175	Ala	His	Arg	Ala	Leu 180
Tyr	Arg	Arg	His	Val 185	Leu	Gly	Ile	Val		Gln	Gly	Pro	Ala	Leu 195
Cys	Phe	Ala	Ala	Ala 200	Ile	Phe	Ser	Leu		Phe	Val	Pro	Leu	Ser 210
Tyr	Leu	Leu	Met	Val 215	Thr	Val	Ile	Leu		Pro	Tyr	Val	Ser	Lys 225
Val	Thr	Gly	Trp	Cys 230	Arg	Asp	Arg	Leu		Gly	His	Arg	Glu	Pro 240
Ser	Ala	His	Pro	Val 245	Glu	Val	Phe	Ser		Asp	Leu	His	Glu	Pro 255
Leu	Ser	Lys	Glu	Arg 260	Val	Glu	Ala	Phe	Ser 265	Asp	Gly	Val	Tyr	Ala 270
Ile	Val	Ala	Thr	Leu 275	Leu	Ile	Leu	Asp	Ile 280	Cys	Glu	Asp	Asn	Val 285
Pro	Asp	Pro	Lys	Asp 290	Val	Lys	Glu	Arg	Phe 295	Ser	Gly	Ser	Leu	Val 300
Ala	Ala	Leu	Ser	Ala 305	Thr	Gly	Pro	Arg	Phe 310	Leu	Ala	Tyr	Phe	Gly 315
Ser	Phe	Ala	Thr	Val 320	Gly	Leu	Leu	Trp	Phe 325	Ala	His	His	Ser	Leu 330
Phe	Leu	His	Val	Arg 335	Lys	Ala	Thr	Arg	Ala 340	Met	Gly	Leu	Leu	Asn 345
Thr	Leu	Ser	Leu	Ala 350	Phe	Val	Gly	Gly	Leu 355	Pro	Leu	Ala	Tyr	Gln 360
Gln	Thr	Ser	Ala	Phe	Ala	Arg	Gln	Pro	Arg 370	Asp	Glu	Leu	Glu	Arg 375
Val	Arg	Val	Ser	Cys 380		Ile	· Ile	Phe	Leu 385	Ala	Ser	Ile	Phe	Gln 390
Leu	Ala	Met	Trp		Thr	Ala	Leu	Let	His 400		Ala	Glu	Thr	Leu 405
			Val	Trp	Phe				415					420
				Leu 425	Tyr				430	1				Ala 435
				Leu 440	Ser				445	)				Leu 450
Met	Gln	ıle	e Ala	Val 455	. Pro	суѕ	s Ala	a Phe	Leu 460	Leu	Leu	a Arç	, Leu	Leu 465
Val	. Gly	, Leu	ı Ala	Leu 470	ı Ala	Thi	Lei	ı Arç		Let	. Arg	g Gl	7 Leu	Ala 480
Arg	g Pro	o Glu	ı His	Pro 485	Pro	Pro	o Ala	a Pro		Gl <sub>y</sub>	glr Glr	n Asp	Asp	Pro 495
Glr	n Ser	Glr	ı Lev		ı Pro	Ala	a Pro	с Суз						

# (2) INFORMATION FOR SEQ ID NO:

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 320 amino acids

143

28:

- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRSTNOT09
  (B) CLONE: 1664634
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:

Met	Ala	Ala	Arg	Leu 5	Asp	Gly	Gly	Phe	Ala 10	Ala	Val	Ser	Arg	Ala 15
Phe	His	Glu	Ile	Arg 20	Ala	Arg	Asn	Pro	Ala 25	Phe	Gln	Pro	Gln	Thr 30
Leu	Met	Asp	Phe	Gly 35	Ser	Gly	Thr	Gly	Ser 40	Val	Thr	Trp	Ala	Ala 45
His	Ser	Ile	Trp	Gly 50	Gln	Ser	Leu	Arg	Glu 55	Tyr	Met	Cys	Val	Asp 60
Arg	Ser	Ala	Ala	Met 65	Leu	Val	Leu	Ala	Glu 70	Lys	Leu	Leu	Thr	Gly 75
Gly	Ser	Glu	Ser	Gly 80	Glu	Pro	Tyr	Ile	Pro 85	Gly	Val	Phe	Phe	Arg 90
Gln	Phe	Leu	Pro	Val 95	Ser	Pro	Lys	Val	Gln 100	Phe	Asp	Val	Val	Val 105
Ser	Ala	Phe	Ser	Leu 110	Ser	Asp	Gln	Leu	Leu 115	Thr	Phe	Ile	Leu	Ser 120
Cys	Asn	Ser	Ser	Leu 125	Leu	His	Ile	Phe	Pro 130	Phe	Cys	Glu	Gln	Val 135
Leu	Val	Glu	Asn	Gly 140	Thr	Lys	Ala	Gly	His 145	Ser	Leu	Leu	Met	Asp 150
	_	Asp		155		_	_	_	160	Lys				165
	_		_	170					175	Pro				180
				185					190	Phe				195
		Ile		200		_		_	205	Pro	_			210
		Met		215				-	220	Pro				225
_	_		_	230					235	Lys	_		_	240
		_		245	_	_		_	250	His				255
				260	-		-	-	265	Gly	_	_	_	270
		_	_	275		_		_	280	Pro	_			285
		Gln		290				_	295	Cys				300
_				305	Ala	Tyr	Ser	Val	Cys 310	Val	Ser	Ser	Ile	Tyr 315
Gly	Ser	Gly	Ser	Leu 320										

- (2) INFORMATION FOR SEQ ID NO: 29:
  - (i) SEQUENCE CHARACTERISTICS:

# RAW SEQUENCE LISTING PATENT APPLICATION US/09/002,485

DATE: 02/07/98 TIME: 14:21:12

INPUT SET: S23256.raw

This Raw Listing contains the General Information Section and up to the first 5 pages.

	1	SEQUENCE LISTING
3	2 3	(1) General Information:
*	3 4	
	5	(i) APPLICANT: Lal, Preeti Hillman, Jennifer L. Corley, Neil C. Cyceler Varl I
•	6	(i) APPLICANT: Lal, Preeti
	7	Hillman, Jennifer L.
	8	Corley, Neil C.
	9	Gdegier, Karr o.
	10	Baugh, Mariah
	11	Sather, Susan Shah, Purvi
	12 13	Sildii, Fulvi
;	14	(ii) TITLE OF INVENTION: HUMAN SIGNAL PEPTIDE-CONTAINING PROTEINS
	15	()
	16	
	17	(iii) NUMBER OF SEQUENCES:154
	18	
	19	(iv) CORRESPONDENCE ADDRESS:
erine See	20 21	(A) ADDRESSEE: INCYTE PHARMACEUTICALS, INC. (B) STREET: 3174 PORTER DRIVE
m	22	(C) CITY: PALO ALTO
un.	23	(D) STATE: CALIFORNIA
#	24	(E) COUNTRY: USA
ļ.ā	25	(F) ZIP: 94304
The state of	26	
	27	( CONDUMED DEADADLE FORM.
i de	28 29	(V) COMPUTER READABLE FORM:  (A) MEDIUM TYPE: Floppy disk
: ::::::::::::::::::::::::::::::::::::	30	(B) COMPUTER: IBM PC compatible
M. C. Francisco	31	(C) OPERATING SYSTEM: PC-DOS/MS-DOS
350	32	(D) SOFTWARE: Word Perfect 6.1 for Windows/MS-DOS 6.2
, del	33	
	34	
	35	(vi) CURRENT APPLICATION DATA:
>	0(36	(A) APPLICATION NUMBER: TO BE ASSIGNED (B) FILING DATE: HEREWITH
	37 38	(C) CLASSIFICATION:
	39	(C) CHABLITONIEN.
	40	
	41	(viii) ATTORNEY/AGENT INFORMATION:
	42	(A) NAME: BILLINGS, LUCY J.
	43	(B) REGISTRATION NUMBER: 36,749
	44	(C) REFERENCE/DOCKET NUMBER: PF-0459 US
	<b>4</b> 5	
	46	

# RAW SEQUENCE LISTING PATENT APPLICATION US/09/002,485

DATE: 02/07/98 TIME: 14:21:16

INPUT SET: S23256.raw

```
47
        (ix) TELECOMMUNICATION INFORMATION:
               (A) TELEPHONE: (650) 855-0555
48
49
               (B) TELEFAX: (650) 845-4166
50
51
52
    (2) INFORMATION FOR SEQ ID NO:
53
54
          (i) SEQUENCE CHARACTERISTICS:
55
56
               (A) LENGTH: 348 amino acids
57
               (B) TYPE: amino acid
58
               (C) STRANDEDNESS: single
59
               (D) TOPOLOGY: linear
60
          (vii) IMMEDIATE SOURCE:
61
62
               (A) LIBRARY: HEARNOT01
63
               (B) CLONE: 305841
64
          (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:
65
66
67
    Met Ala Ala Thr Leu Gly Pro Leu Gly Ser Trp Gln Gln Trp Arg
68
69
    Arg Cys Leu Ser Ala Arg Asp Gly Ser Arg Met Leu Leu Leu
70
                                           25
    Leu Leu Cly Ser Cly Gln Cly Pro Gln Gln Val Gly Ala Gly
71
72
                                           40
    Gln Thr Phe Glu Tyr Leu Lys Arg Glu His Ser Leu Ser Lys Pro
73
74
                      50
                                           55
75
    Tyr Gln Gly Val Gly Thr Gly Ser Ser Leu Trp Asn Leu Met
76
                                           70
                                                               75
                      65
    Gly Asn Ala Met Val Met Thr Gln Tyr Ile Arg Leu Thr Pro Asp
77
78
                      80
                                           85
79
    Met Gln Ser Lys Gln Gly Ala Leu Trp Asn Arg Val Pro Cys Phe
80
                      95
                                          100
81
    Leu Arg Asp Trp Glu Leu Gln Val His Phe Lys Ile His Gly Gln
82
                     110
                                          115
83
    Gly Lys Lys Asn Leu His Gly Asp Gly Leu Ala Ile Trp Tyr Thr
84
                     125
                                          130
85
    Lys Asp Arg Met Gln Pro Gly Pro Val Phe Gly Asn Met Asp Lys
86
                     140
                                          145
    Phe Val Gly Leu Gly Val Phe Val Asp Thr Tyr Pro Asn Glu Glu
87
88
                     155
                                         160
    Lys Gln Gln Glu Arg Val Phe Pro Tyr Ile Ser Ala Met Val Asn
89
90
                     170
                                          175
91
    Asn Gly Ser Leu Ser Tyr Asp His Glu Arg Asp Gly Arg Pro Thr
92
                     185
                                          190
93
    Glu Leu Gly Gly Cys Thr Ala Ile Val Arg Asn Leu His Tyr Asp
94
                     200
                                          205
95
    Thr Phe Leu Val Ile Arg Tyr Val Lys Arg His Leu Thr Ile Met
96
                     215
                                          220
97
    Met Asp Ile Asp Gly Lys His Glu Trp Arg Asp Cys Ile Glu Val
98
                     230
                                          235
99
    Pro Gly Val Arg Leu Pro Arg Gly Tyr Tyr Phe Gly Thr Ser Ser
```

the transfer of the transfer o





RAW SEQUENCE LISTING PATENT APPLICATION US/09/002,485 DATE: 02/07/98 TIME: 14:21:19

														IN	PUT SET: S23256.raw
100					245					250					255
101	Ile	Thr	Gly	Asp	Leu	Ser	Asp	Asn	His	Asp	Val	Ile	Ser	Leu	Lys
102			-	_	260		_			265					270
103	Leu	Phe	Glu	Leu	Thr	Val	Glu	Arg	Thr	Pro	Glu	Glu	Glu	Lys	Leu
104					275					280					285
105	His	Arg	Asp	Val	Phe	Leu	${\tt Pro}$	Ser	Val	Asp	Asn	Met	Lys	Leu	Pro
106					290					295					300
107	Glu	Met	Thr	Ala	Pro	Leu	Pro	Pro	Leu	Ser	Gly	Leu	Ala	Leu	Phe
108					305					310					315
109	Leu	Ile	Val	Phe	Phe	Ser	Leu	Val	Phe	Ser	Val	Phe	Ala	Ile	
110					320					325		_			330
111	Ile	Gly	Ile	Ile	Leu	$\mathtt{Tyr}$	Asn	Lys	Trp		Glu	Gln	Ser	Arg	
112					335					340					345
113	Arg	Phe	Tyr												
114															
115															
116															
117		T 3.7 EV	ND1481	T-031	EOD	OHO.	TD 1			2.					
118	(2)	INE	JKMA'	LTON	FOR	SEQ	י עד	NO:		2:					
119		/ -	C CT	<b>77 773 37</b>	מס פור	13 D 3 (	ameio.	TOMT	<b>70.</b>						
120		( τ			CE CI ENGTI					a.					
121 122			-		YPE:				acı	20					
123				-	rani				ale						
124			•	•	OPOLO				9-0						
125			١.	, -	J. J.										
126		(v:	ii) :	EMME:	DIATE	E SOI	JRCE	:							
127		,			IBRAI				2						
128			•	,	LONE										
129			•	•											
130		(x:	i) Si	EQUE	NCE I	DESC	RIPT	ION:	SEQ	ID I	OF:	2:			
131															
132	Met	Gly	Met	Ser	Ser	Leu	Lys	Leu	Leu	Lys	Tyr	Val	Leu	Phe	Phe
133					5					10				_	15
134	Phe	Asn	Leu	Leu	Phe	Trp	Ile	Cys	Gly		Cys	Ile	Leu	Gly	
135	_	_			20	_				25			_	_,	30
136	Gly	Ile	Tyr	Leu	Leu	Ile	His	Asn	Asn	_	GTA	Val	Leu	Phe	
137		_	_		_ 35		_	7		40	<b></b> 1	•• - T	-1-	7	45
138	Asn	Leu	Pro	Ser	Leu	'l'nr	Leu	СТÄ	Asn		Pne	va⊥	тте	vaı	
139		-7 -	<b>-</b> 3 -		50	**-7		D1	T	55	a	1404	a1	Com	60
140	ser	тте	тте	мет		val	ATA	Pne	Leu		Cys	Mec	сту	ser	75
141	T	a1	2 ~ ~	T	65 Cys	T 011	T 011	Wot	802	70 Bho	Dho	Tla	LOII	Lou	
142 143	гуѕ	GIU	ASII	гур	80	ьeu	ьeu	Met	Ser	85	FILE	TTE	цец	пец	90
	т1.	Tla	T 011	T 011	Ala	@lu	Te V	Thr	T.011		ΤlΔ	T. 🗕 11	T.011	Phe	
144 145	TTE	TTE	neu	пeп	95	JIU	* a T	T11T	มสน	100		u	Luu	- 116	105
145	ጥ፣ታጉ	Glu	g] n	T.17e	Leu	Δsn	Glu	ጥነንዮ	Val		Lvs	Gl v	Leu	Thr	
147	- y -	OIU	O-111	цуз	110	A DII	O.L.u	+ Y +	+ u <b>1</b>	115	- 75	~-1			120
148	Ser	Ile	His	Ara	Tyr	His	Ser	Asp	Asn		Thr	Lvs	Ala	Ala	
149	~~-			9	125					130		-1-			135
150	Asp	Ser	Ile	Gln	Ser	Phe	Leu	Gln	Cys		Gly	Ile	Asn	Gly	
151	<b>L</b>				140				<b>4</b> -	145	-			-	150
152	Ser	Asp	Leu	Asp		Gly	Ser	Pro	Ala	Ser	Cys	Pro	Ser	Asp	Arg
		-		-		-					-			_	=

18.

# PAGE: 4 RAW SEQUENCE LISTING PATENT APPLICATION US/09/002,485

DATE: 02/07/98 TIME: 14:21:23

# INPUT SET: S23256.raw

```
160
                                                              165
153
                     155
154
     Lys Val Glu Gly Cys Tyr Ala Lys Glu Asp Phe Gly Phe Ile Gln
155
                     170
                                          175
     Phe Pro Val Tyr Arg Asn His His Leu Cys Met Cys Asp
156
                                          190
157
                      185
158
159
160
161
```

3:

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 342 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BEPINOT01
- (B) CLONE: 546656

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:

Met Ser Leu His Gly Lys Arg Lys Glu Ile Tyr Lys Tyr Glu Ala

177	Mec	Der	пеа	III	5	БУБ	n. g	пуы	OLU	10	- 7 -	טעם	- y -	Olu	15
178 179	Pro	Trp	Thr	Val	Tyr 20	Ala	Met	Asn	Trp	Ser 25	Val	Arg	Pro	Asp	Lys 30
180 181	Arg	Phe	Arg	Leu	Ala 35	Leu	Gly	Ser	Phe	Val 40	Glu	Glu	Tyr	Asn	Asn 45
182 183	Lys	Val	Gln	Leu	Val 50	Gly	Leu	Asp	Glu	Glu 55	Ser	Ser	Glu	Phe	Ile 60
184 185	Cys	Arg	Asn	Thr	Phe 65	Asp	His	Pro	Tyr	Pro 70	Thr	Thr	Lys	Leu	Met 75
186 187	Trp	Ile	Pro	Asp	Thr 80	Lys	Gly	Val	Tyr	Pro 85	Asp	Leu	Leu	Ala	Thr 90
188 189	Ser	Gly	Asp	Tyr	Leu 95	Arg	Val	Trp	Arg	Val 100	Gly	Glu	Thr	Glu	Thr 105
190 191	Arg	Leu	Glu	Cys	Leu 110	Leu	Asn	Asn	Asn	Lys 115	Asn	Ser	Asp	Phe	Cys 120
192 193	Ala	Pro	Leu	Thr	Ser 125	Phe	Asp	Trp	Asn	Glu 130	Val	Asp	Pro	Tyr	Leu 135
194 195	Leu	Gly	Thr	Ser	Ser 140	Ile	Asp	Thr	Thr	Cys 145	Thr	Ile	Trp	Gly	Leu 150
196 197	Glu	Thr	Gly	Gln	Val 155	Leu	Gly	Arg	Val	Asn 160	Leu	Val	Ser	Gly	His 165
198 199	Val	Lys	Thr	Gln	Leu 170	Ile	Ala	His	Asp	Lys 175	Glu	Val	Tyr	Asp	Ile 180
200 201	Ala	Phe	Ser	Arg	Ala 185	Gly	Gly	Gly	Arg	Asp 190	Met	Phe	Ala	Ser	Val 195
202 203	Gly	Ala	Asp	Gly	Ser 200	Val	Arg	Met	Phe	Asp 205	Leu	Arg	His	Leu	Glu 210
204 205	His	Ser	Thr	Ile	Ile 215	Tyr	Glu	Asp	Pro	Gln 220	His	His	Pro	Leu	Leu 225





# **RAW SEQUENCE LISTING**PATENT APPLICATION *US/09/002*,485 DATE: 02/07/98 TIME: 14:21:27

														IN	PUT	SET: S	23256.	raw
206	Ara	Leu	Cvs	Trp	Asn	Lys	Gln	Asp	Pro	Asn	Tyr	Leu	Ala					- •••
207	5			-	230	-		-		235	-				240			
208	Ala	Met	Asp	Gly	Met	Glu	Val	Val	Ile	Leu	Asp	Val	Arg	Val	Pro			
209					245					250					255			
210	Cys	Thr	Pro	Val	Ala	Arg	Leu	Asn	Asn		Arg	Ala	Cys	Val				
211	_	_	_		260				_	265			_	1	270			
212	Gly	Ile	Ala	Trp		Pro	His	Ser	Ser		His	Ile	Cys	Thr				
213	. 7		3	TT2	275		T	T1.	Пии	280	т1.	al n	aln	Wot	285 Bro			
214	Ата	Asp	Asp	HIS	290	Ата	rea	тте	тгр	295	TTG	GIH	GTII	мес	300			
215 216	X ~~	Ala	T1^	alu		Dro	т1а	T. 611	λla		ሞክ r	Δla	Glu	G] v				
216	Arg	АТа	TTE	GIU	305	PIU	TTG	ьeu	АТа	310	1111	лта	GLU	OLY	315			
217	Tle	Asn	Δsn	Va1		Trp	Ala	Ser	Thr	-	Pro	Asp	Tro	Ile				
219		11011		,	320					325		<u>-</u>			330			
220	Ile	Cys	Tyr	Asn		Cys	Leu	Glu	Ile		Arg	Val						
221		-	-		335	•				340	_							
222																		
223																		
224																		
225																		
226	(2)	INF	ORMA'	rion	FOR	SEQ	ID 1	NO:		4:								
227																		
228		(i	•	-	CE CI					-								
229			•	•	ENGTI				acı	ds								
230			•	•	YPE:				T									
231					rrani				эте									
232 233			( :	יז (ט	OPOL	JGI:	T. T. I. I.	ear										
234		/ 57		TMME	DIATI	7 SOI	TRCE	•										
235		( v .			IBRAI				3									
236					LONE				-									
237			``	-,														
238		( x.	i) S	EQUE:	NCE I	DESC	RIPT	ION:	SEQ	ID 1	NO:	4 :						
239		•	·															
240	Met	Glu	Glu	Leu	Asp	Gly	Glu	Pro	Thr	Val	Thr	Leu	Ile	Pro	Gly			
241					5					10					15			
242	Val	Asn	Ser	Lys	Lys	Asn	Gln	Met	Tyr	Phe	Asp	Trp	Gly	Pro				
243					20	_	_		_	25		_		_	30			
244	Glu	Met	Leu	Val		Glu	Thr	Ser	Phe		Lys	Lys	Glu	Lys				
245				_	35	_	_	_,		40	<b>-</b> -7			<b>.</b>	45			
246	Glu	Met	Val	Pro		Cys	Pro	Phe	TTe		тте	тте	arg	гàг				
247		3	17-7	m	50	<b>~1</b>	т1	т	7 m~	55	Τ	Dho	λ <b>~</b> ~	<b>01</b> 11	60			
248	vaı	Asp	val	ryr		GIN	тте	Leu	Arg	_	ьeu	PHE	ASII	GIU	75			
249					65	_		_		70		_		_	, 3			

His Gly Ile Phe Leu Gly Leu Gln Arg Ile Asp Glu Glu Leu Thr

Gly Lys Ser Arg Lys Ser Gln Leu Val Arg Val Ser Lys Asn Tyr

Arg Ser Val Ile Arg Ala Cys Met Glu Glu Met His Gln Val Ala

Ile Ala Ala Lys Asp Pro Ala Asn Gly Arg Gln Phe Ser Ser Gln

Val Ser Ile Leu Ser Ala Met Glu Leu Ile Trp Asn Leu Cys Glu

PAGE: 1

# SEQUENCE VERIFICATION REPORT PATENT APPLICATION US/09/002,485

DATE: 02/07/98 TIME: 14:21:31

INPUT SET: S23256.raw

Line

Error

Original Text

36

Wrong application Serial Number

(A) APPLICATION NUMBER: TO BE ASSIGNED

- (A) LENGTH: 117 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: PROSTUT10
- (B) CLONE: 1690990

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:

Met Asp Asn Lys Gly Ile Tyr Pro Gly Ala Val Phe Tyr His Asp 15 Ser Phe Thr Glu Ser Arg Val Val Leu Leu Arg Ile Arg Thr Leu 20 25 30 Val Pro Tyr Ser Pro Pro Asp Cys Pro Thr Thr Thr Ala Tyr 35 40 45 Ser Pro Phe Pro Asn His Gly Gln Gln Ile Glu Leu Leu Thr Glu 50 55 Val Ser Phe Arg Trp Ile Ser Gln Pro Phe Pro His Arg Pro His 65 70 Arg Glu Thr Val Thr Asp Cys Tyr Ser Pro Asn Thr Gln Val Lys 80 85 90 Ser Asn Ala Gly Arg Asn Asn Ser Lys Ser Phe Asn Phe Leu Ile 95 100 Leu Leu Lys Ile Leu Thr Glu Ala Ser Arg Phe 110

30:

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 298 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: DUODNOT02
- (B) CLONE: 1704050

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:

Met Ala Arg Arg Ser Arg His Arg Leu Leu Leu Leu Leu Arg 10 Tyr Leu Val Val Ala Leu Gly Tyr His Lys Ala Tyr Gly Phe Ser 20 25 Ala Pro Lys Asp Gln Gln Val Val Thr Ala Val Glu Tyr Gln Glu 35 40 Ala Ile Leu Ala Cys Lys Thr Pro Lys Lys Thr Val Ser Ser Arg 50 55 60 Leu Glu Trp Lys Lys Leu Gly Arg Ser Val Ser Phe Val Tyr Tyr 65 70 Gln Gln Thr Leu Gln Gly Asp Phe Lys Asn Arg Ala Glu Met Ile 85 90 Asp Phe Asn Ile Arg Ile Lys Asn Val Thr Arg Ser Asp Ala Gly 95 100 Lys Tyr Arg Cys Glu Val Ser Ala Pro Ser Glu Gln Gly Gln Asn 110 115 120

```
Leu Glu Glu Asp Thr Val Thr Leu Glu Val Leu Val Ala Pro Ala
                125
                                   130
Val Pro Ser Cys Glu Val Pro Ser Ser Ala Leu Ser Gly Thr Val
                140
                                    145
Val Glu Leu Arg Cys Gln Asp Lys Glu Gly Asn Pro Ala Pro Glu
                155
                                    160
Tyr Thr Trp Phe Lys Asp Gly Ile Arg Leu Leu Glu Asn Pro Arg
                170
                                    175
Leu Gly Ser Gln Ser Thr Asn Ser Ser Tyr Thr Met Asn Thr Lys
                185
                                    190
                                                        195
Thr Gly Thr Leu Gln Phe Asn Thr Val Ser Lys Leu Asp Thr Gly
                200
                                    205
                                                        210
Glu Tyr Ser Cys Glu Ala Arg Asn Ser Val Gly Tyr Arg Arg Cys
                215
                                    220
Pro Gly Lys Arg Met Gln Val Asp Asp Leu Asn Ile Ser Gly Ile
                230
                                    235
Ile Ala Ala Val Val Val Ala Leu Val Ile Ser Val Cys Gly
                245
                                    250
Leu Gly Val Cys Tyr Ala Gln Arg Lys Gly Tyr Phe Ser Lys Glu
                260
                                    265
                                                        270
Thr Ser Phe Gln Lys Ser Asn Ser Ser Ser Lys Ala Thr Thr Met
                275
                                    280
Ser Glu Asn Asp Phe Lys His Thr Lys Ser Phe Ile Ile
                290
                                    295
```

### (2) INFORMATION FOR SEQ ID NO:

31:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 118 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: PROSNOT16
- (B) CLONE: 1711840

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:

```
Met Gln His Arg Gly Phe Leu Leu Thr Leu Leu Ala Leu Leu
                                     10
Ala Leu Thr Ser Ala Val Ala Lys Lys Gln Asp Lys Val Lys
                 20
                                     25
                                                         30
Gly Gly Pro Gly Ser Glu Cys Ala Glu Trp Ala Trp Gly Pro Cys
                 35
                                     40
Thr Pro Ser Ser Lys Gly Phe Ala Ala Val Gly Phe Pro Arg Gly
                 50
                                     55
                                                         60
Pro Pro Trp Gly Gly Pro Arg Thr Gln Pro Ala Val Leu Val Glu
                 65
                                     70
Arg Val Ala Pro Gly Lys Leu Glu Arg Lys Glu Phe Trp Ala Pro
                 80
                                     85
Gly Leu Trp Lys Val Gly Gln Ile Phe Trp Lys Lys Thr Trp Arg
                                    100
Val Cys Arg Ser Val Lys Trp Gly Arg Gly Gln Lys Asn
                110
                                    115
```

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 248 amino acids

  - (B) TYPE: amino acid (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:

Met	Gln	Thr	Cys	Pro 5	Leu	Ala	Phe	Pro	Gly 10	His	Val	Ser	Gln	Ala 15
Leu	Gly	Thr	Leu	Leu 20	Phe	Leu	Ala	Ala	Ser 25	Leu	Ser	Ala	Gln	Asn 30
Glu	Gly	Trp	Asp	Ser 35	Pro	Ile	Суѕ	Thr	Glu 40	Gly	Val	Val	Ser	Val 45
Ser	Trp	Gly	Glu	Asn 50	Thr	Val	Met	Ser	Cys 55		Ile	Ser	Asn	Ala 60
Phe	Ser	His	Val	Asn 65	Ile	Lys	Leu	Arg	Ala 70	His	Gly	Gln	Glu	Ser 75

Ala	Ile	Phe	Asn	Glu 80	Val	Ala	Pro	Gly	Tyr 85	Phe	Ser	Arg	Asp	Gly 90
Trp	Gln	Leu	Gln	Val 95	Gln	Gly	Gly	Val	Ala 100	Gln	Leu	Val	Ile	Lys 105
Gly	Ala	Arg	Asp	Ser 110	His	Ala	Gly	Leu	Tyr 115	Met	Trp	His	Leu	Val 120
				125		Arg			130					135
Ala	Glu	Pro	Gln	Ser 140	Ala	Pro	Asp	Thr	Gly 145	Phe	Trp	Pro	Val	Pro 150
Ala	Val	Val	Thr	Ala 155	Val	Phe	Ile	Leu	Leu 160	Val	Ala	Leu	Val	Met 165
Phe	Ala	Trp	Tyr	Arg 170	Cys	Arg	Cys	Ser	Gln 175	Gln	Arg	Arg	Glu	Lys 180
Lys	Phe	Phe	Leu	Leu 185	Glu	Pro	Gln	Met	Lys 190	Val	Ala	Ala	Leu	Arg 195
Ala	Gly	Ala	Gln	Gln 200	Gly	Leu	Ser	Arg	Ala 205	Ser	Ala	Glu	Leu	Trp 210
Thr	Pro	Asp	Ser	Glu 215	Pro	Thr	Pro	Arg	Pro 220	Leu	Ala	Leu	Val	Phe 225
Lys	Pro	Ser	Pro	Leu 230	Gly	Ala	Leu	Glu	Leu 235	Leu	Ser	Pro	Gln	Pro 240
Leu	Phe	Pro	Tyr		Ala	Asp	Pro							

#### (2) INFORMATION FOR SEQ ID NO: 33:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 150 amino acids (B) TYPE: amino acid

  - (C) STRANDEDNESS: single

(D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

(A) LIBRARY: STOMTUT02

(B) CLONE: 1750632

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:

Met Leu Glu Glu Gly Ser Phe Arg Gly Arg Thr Ala Asp Phe Val 10 Phe Met Phe Leu Phe Gly Gly Val Leu Met Thr Val Ser Phe Pro 20 25 Gln Ala Leu Glu Pro Arg Ala Arg Ala Pro Arg Arg Pro Ala Cys 35 40 Val Gly Pro Gly Ala Asn Thr Ala Met Pro Glu Arg Asp Thr Val 50 55 Ala Val Ser Ser Leu Ala Pro Phe Leu Pro Trp Ala Leu Met Gly 65 70 Phe Ser Leu Leu Gly Asn Ser Ile Leu Val Asp Leu Leu Gly 85 80 Ile Ala Val Gly His Ile Tyr Tyr Phe Leu Glu Asp Val Phe Pro 95 100 Asn Gln Pro Gly Gly Lys Arg Leu Leu Gln Thr Pro Gly Phe Leu 110 120 115 Lys Leu Leu Asp Ala Pro Ala Glu Asp Pro Asn Tyr Leu Pro 125 130 Leu Pro Glu Glu Gln Pro Gly Pro His Leu Pro Pro Pro Gln Gln 140 145

### (2) INFORMATION FOR SEQ ID NO:

### (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 431 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:

Met	Trp	Ala	Leu	Gly 5		Ala	Gly	Phe	Ala 10	Asn	Leu	Thr	Glu	Gly 15
Leu	Lys	Val	Trp	Leu 20	Gly	Ile	Met	Leu	Pro 25	Val	Leu	Gly	Ile	Lys 30
Ser	Leu	Ser	Pro	Phe 35		Ile	Thr	Tyr	Leu 40	Asp	Arg	Leu	Leu	Leu 45
Met	His	Pro	Asn	Leu 50	Thr	Lys	Gly	Phe	Gly 55	Met	Ile	Gly	Pro	Lys 60
Asp	Phe	Phe	Pro	Leu 65	Leu	Asp	Phe	Ala	Tyr 70	Met	Pro	Asn	Asn	Ser 75
Leu	Thr	Pro	Ser	Leu 80	Gln	Glu	Gln	Leu	Cys 85	Gln	Leu	Tyr	Pro	Arg 90
Leu	Lys	Met	Leu	Ala 95	Phe	Gly	Ala	Lys	Pro 100	Asp	Ser	Thr	Leu	His 105
Thr	Tyr	Phe	Pro	Ser 110				Arg		Thr	Pro	Ser	Cys	Pro 120

Pro	Glu	Met	Lys	Lys 125	Glu	Leu	Leu	Ser	Ser 130	Leu	Thr	Glu	Cys	Leu 135
Thr	Val	Asp	Pro	Leu 140	Ser	Ala	Ser	Val	Trp 145	Arg	Gln	Leu	Tyr	Pro 150
Lys	His	Leu	Ser	Gln 155	Ser	Ser	Leu	Leu	Leu 160	Glu	His	Leu	Leu	Ser 165
Ser	Trp	Glu	Gln	Ile 170	Pro	Lys	Lys	Val	Gln 175	Lys	Ser	Leu	Gln	Glu 180
Thr	Ile	Gln	Ser	Leu 185	Lys	Leu	Thr	Asn	Gln 190	Glu	Leu	Leu	Arg	Lys 195
Gly	Ser	Ser	Asn	Asn 200	Gln	Asp	Val	Val	Thr 205	Cys	Asp	Met	Ala	Cys 210
Lys	Gly	Leu	Leu	Gln 215	Gln	Val	Gln	Gly	Pro 220	Arg	Leu	Pro	Trp	Thr 225
Arg	Leu	Leu	Leu	Leu 230	Leu	Leu	Val	Phe	Ala 235	Val	Gly	Phe	Leu	Cys 240
His	Asp	Leu	Arg	Ser 245	His	Ser	Ser	Phe	Gln 250	Ala	Ser	Leu	Thr	Gly 255
Arg	Leu	Leu	Arg	Ser 260	Ser	Gly	Phe	Leu	Pro 265	Ala	Ser	Gln	Gln	Ala 270
Cys	Ala	Lys	Leu	Tyr 275	Ser	Tyr	Ser	Leu	Gln 280	Gly	Tyr	Ser	Trp	Leu 285
Gly	Glu	Thr	Leu	Pro 290	Leu	Trp	Gly	Ser	His 295	Leu	Leu	Thr	Val	Val 300
Arg	Pro	Ser	Leu	Gln 305	Leu	Ala	Trp	Ala	His 310	Thr	Asn	Ala	Thr	Val 315
Ser	Phe	Leu	Ser		His	Cys	Ala	Ser	His 325	Leu	Ala	Trp	Phe	Gly 330
Asp	Ser	Leu	Thr	Ser 335	Leu	Ser	Gln	Arg	Leu 340	Gln	Ile	Gln	Leu	Pro 345
Asp	Ser	Val	Asn	Gln 350	Leu	Leu	Arg	Tyr	Leu 355	Arg	Glu	Leu	Pro	Leu 360
Leu	Phe	His	Gln	Asn 365	Val	Leu	Leu	Pro	Leu 370	Trp	His	Leu	Leu	Leu 375
Glu	Ala	Leu	Ala	Trp 380	Ala	Gln	Glu	His	Cys 385	His	Glu	Ala	Cys	Arg 390
Gly	Glu	Val	Thr	Trp 395	Asp	Cys	Met	Lys	Thr 400	Gln	Leu	Ser	Glu	Ala 405
Val	His	Trp	Thr	Trp 410	Leu	Cys	Leu	Gln	Asp 415	Ile	Thr	Val	Ala	Phe 420
Leu	Asp	Trp	Ala	Leu 425	Ala	Leu	Ile	Ser	Gln 430	Gln				

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 278 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: PROSNOT20
  - (B) CLONE: 1818761

35:

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:

Met Gln Trp Leu Arg Val Arg Glu Ser Pro Gly Glu Ala Thr Gly 10 His Arg Val Thr Met Gly Thr Ala Ala Leu Gly Pro Val Trp Ala 20 25 Ala Leu Leu Phe Leu Leu Met Cys Glu Ile Pro Met Val Glu 35 40 Leu Thr Phe Asp Arg Ala Val Ala Ser Gly Cys Gln Arg Cys Cys 50 55 Asp Ser Glu Asp Pro Leu Asp Pro Ala His Val Ser Ser Ala Ser 70 65 Ser Ser Gly Arg Pro His Ala Leu Pro Glu Ile Arg Pro Tyr Ile 85 80 Asn Ile Thr Ile Leu Lys Gly Asp Lys Gly Asp Pro Gly Pro Met 100 95 Gly Leu Pro Gly Tyr Met Gly Arg Glu Gly Pro Gln Gly Glu Pro 110 115 Gly Pro Gln Gly Ser Lys Gly Asp Lys Gly Glu Met Gly Ser Pro 130 125 135 Gly Ala Pro Cys Gln Lys Arg Phe Phe Ala Phe Ser Val Gly Arg 140 145 Lys Thr Ala Leu His Ser Gly Glu Asp Phe Gln Thr Leu Leu Phe 155 160 Glu Arg Val Phe Val Asn Leu Asp Gly Cys Phe Asp Met Ala Thr 170 175 Gly Gln Phe Ala Ala Pro Leu Arg Gly Ile Tyr Phe Phe Ser Leu 185 190 Asn Val His Ser Trp Asn Tyr Lys Glu Thr Tyr Val His Ile Met 200 205 His Asn Gln Lys Glu Ala Val Ile Leu Tyr Ala Gln Pro Ser Glu 215 220 225 Arg Ser Ile Met Gln Ser Gln Ser Val Met Leu Asp Leu Ala Tyr 230 235 Gly Asp Arg Val Trp Val Arg Leu Phe Lys Arg Gln Arg Glu Asn 245 250 Ala Ile Tyr Ser Asn Asp Phe Asp Thr Tyr Ile Thr Phe Ser Gly 260 265 His Leu Ile Lys Ala Glu Asp Asp 275

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 286 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: GBLATUT01
- (B) CLONE: 1824469
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

Met Glu Glu Lys Arg Arg Arg Ala Arg Val Gln Gly Ala Trp Ala 5 10 15 Ala Pro Val Lys Ser Gln Ala Ile Ala Gln Pro Ala Thr Thr Ala

```
20
Lys Ser His Leu His Gln Lys Pro Gly Gln Thr Trp Lys Asn Lys
                 35
                                     40
Glu His His Leu Ser Asp Arg Glu Phe Val Phe Lys Glu Pro Gln
                 50
                                      55
Gln Val Val Arg Arg Ala Pro Glu Pro Arg Val Ile Asp Arg Glu
                                     70
                                                          75
                 65
Gly Val Tyr Glu Ile Ser Leu Ser Pro Thr Gly Val Ser Arg Val
                 80
                                                          90
Cys Leu Tyr Pro Gly Phe Val Asp Val Lys Glu Ala Asp Trp Ile
                 95
                                     100
                                                         105
Leu Glu Gln Leu Cys Gln Asp Val Pro Trp Lys Gln Arg Thr Gly
                                     115
                110
Ile Arg Glu Asp Ile Thr Tyr Gln Gln Pro Arg Leu Thr Ala Trp
                                    130
                                                         135
                125
Tyr Gly Glu Leu Pro Tyr Thr Tyr Ser Arg Ile Thr Met Glu Pro
                140
                                     145
Asn Pro His Trp His Pro Val Leu Arg Thr Leu Lys Asn Arg Ile
                155
                                     160
Glu Glu Asn Thr Gly His Thr Phe Asn Ser Leu Leu Cys Asn Leu
                                     175
                170
Tyr Arg Asn Glu Lys Asp Ser Val Asp Trp His Ser Asp Asp Glu
                185
                                     190
                                                         195
Pro Ser Leu Gly Arg Cys Pro Ile Ile Ala Ser Leu Ser Phe Gly
                200
                                     205
                                                         210
Ala Thr Arg Thr Phe Glu Met Arg Lys Lys Pro Pro Pro Glu Glu
                215
                                     220
Asn Gly Asp Tyr Thr Tyr Val Glu Arg Val Lys Ile Pro Leu Asp
                230
                                     235
His Gly Thr Leu Leu Ile Met Glu Gly Ala Thr Gln Ala Asp Trp
                245
                                     250
Gln His Arg Val Pro Lys Glu Tyr His Ser Arg Glu Pro Arg Val
                260
                                     265
Asn Leu Thr Phe Arg Thr Val Tyr Pro Asp Pro Arg Gly Ala Pro
                275
                                     280
                                                         285
Trp
```

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 404 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: PROSNOT19
- (B) CLONE: 1864292

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 37:

Met Lys Met Glu Glu Ala Val Gly Lys Val Glu Glu Leu Ile Glu S 10 10 15 Ser Glu Ala Pro Pro Lys Ala Ser Glu Glu Glu Thr Ala Lys Glu 20 25 30 Glu Asp Gly Ser Val Glu Leu Glu Ser Gln Val Gln Lys Asp Gly 35

```
Val Ala Asp Ser Thr Val Ile Ser Ser Met Pro Cys Leu Leu Met
                 50
                                     55
Glu Leu Arg Arg Asp Ser Ser Glu Ser Gln Leu Ala Ser Thr Glu
                                     70
                 65
Ser Asp Lys Pro Thr Thr Gly Arg Val Tyr Glu Ser Asp Pro Ser
                                     85
                 80
Asn His Cys Met Leu Ser Pro Ser Ser Ser Gly His Leu Ala Asp
                                    100
                 95
Ser Asp Thr Leu Ser Ser Ala Glu Glu Asn Glu Pro Ser Gln Ala
                                    115
                110
Glu Thr Ala Val Glu Gly Asp Pro Ser Gly Val Ser Gly Ala Thr
                                    130
                125
Val Gly Arg Lys Ser Arg Arg Ser Arg Ser Glu Ser Glu Thr Ser
                                    145
                140
Thr Met Ala Ala Lys Lys Asn Arg Gln Ser Ser Asp Lys Gln Asn
                                    160
                155
Gly Arg Val Ala Lys Val Lys Gly His Arg Ser Gln Lys His Lys
                170
                                    175
Glu Arg Ile Arg Leu Leu Arg Gln Lys Arg Glu Ala Ala Arg
                185
                                    190
Lys Lys Tyr Asn Leu Leu Gln Asp Ser Ser Thr Ser Asp Ser Asp
                                    205
                200
Leu Thr Cys Asp Ser Ser Thr Ser Ser Ser Asp Asp Glu Glu
                215
                                     220
Val Ser Gly Ser Ser Lys Thr Ile Thr Ala Glu Ile Pro Asp Gly
                                     235
                230
Pro Pro Val Val Ala His Tyr Asp Met Ser Asp Thr Asn Ser Asp
                                     250
                245
Pro Glu Val Val Asn Val Asp Asn Leu Leu Ala Ala Val Val
                                     265
                260
Gln Glu His Ser Asn Ser Val Gly Gln Asp Thr Gly Ala Thr
                275
                                     280
Trp Arg Thr Ser Gly Leu Leu Glu Glu Leu Asn Ala Glu Ala Gly
                290
                                     295
                                                         300
His Leu Asp Pro Gly Phe Leu Ala Ser Asp Lys Thr Ser Ala Gly
                                     310
                                                         315
                305
Asn Ala Pro Leu Asn Glu Glu Ile Asn Ile Ala Ser Ser Asp Ser
                320
                                     325
Glu Val Glu Ile Val Gly Val Gln Glu His Ala Arg Cys Val His
                335
                                     340
                                                         345
Pro Arg Gly Gly Val Ile Gln Ser Val Ser Ser Trp Lys His Gly
                                                         360
                350
                                     355
Ser Gly Thr Gln Tyr Val Ser Thr Arg Gln Thr Gln Ser Trp Thr
                                     370
                365
Ala Val Thr Pro Gln Gln Thr Trp Ala Ser Pro Ala Glu Val Val
                                     385
                380
Asp Leu Thr Leu Asp Glu Asp Ser Arg Arg Lys Tyr Leu Leu
                395
                                     400
```

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 405 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

(A) LIBRARY: THP1NOT01
(B) CLONE: 1866437

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 38:

Met	Phe	Val	Gln	Glu 5	Glu	Lys	Ile	Phe	Ala 10	Gly	Lys	Val	Leu	Arg 15
Leu	His	Ile	Cys		Ser	Asp	Gly	Ala		Trp	Leu	Glu	Glu	Ala 30
Thr	Glu	Asp	Thr		Val	Glu	Lys	Leu		Glu	Arg	Cys	Leu	
His	Cys	Ala	His		Ser	Leu	Glu	Asp		Lys	Ser	Ile	Thr	His 60
His	Lys	Leu	Ile		Ala	Ala	Ser	Glu		Val	Leu	Ser	Asp	Ala 75
Arg	Thr	Ile	Leu		Glu	Asn	Ile	Gln		Gln	Asp	Val	Leu	Leu 90
Leu	Lys	Lys	Lys		Ala	Pro	Ser	Pro		Pro	Lys	Met	Ala	Asp 105
Val	Ser	Ala	Glu		Lys	Lys	Lys	Gln	Asp 115	Gln	Lys	Ala	Pro	Asp 120
Lys	Glu	Ala	Ile		Arg	Ala	Thr	Ala	Asn 130	Leu	Pro	Ser	Tyr	Asn 135
Met	Asp	Arg	Ala		Val	Gln	Thr	Asn	Met 145	Arg	Asp	Phe	Gln	Thr 150
Glu	Leu	Arg	Lys		Leu	Val	Ser	Leu	Ile 160	Glu	Val	Ala	Gln	Lys 165
				170	Pro				175					180
				185	Glu				190					195
				200	Thr				205					210
	_			215	Leu				220					225
	_			230					235					240
				245	Pro				250					255
				260					265					270
_	_			275					280					285
				290					295					300
_		_		305					310					Asn 315
				320					325					Lys 330
				335					340					Pro 345
				350					355					Leu 360
				365					370					Asn 375
				380	)				385					Pro 390
Val	Met	Leu	. Gln	. Il∈ 395		Arg	Ile	Phe	Gln 400		Leu	ı Asr	Arg	Thr 405

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 177 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: SKINBIT01
  - (B) CLONE: 1871375
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:

Met	Val	Met	His	Asn 5	Ser	Asp	Pro	Asn	Leu 10	His	Leu	Leu	Ala	Glu 15
Gly	Ala	Pro	Ile	Asp 20	Trp	Gly	Glu	Glu	Tyr 25	Ser	Asn	Ser	Gly	Gly 30
Gly	Gly	Ser	Pro	Ala 35	Pro	Ala	Pro	Arg	Ser 40	Gln	Pro	Pro	Ser	Arg 45
Lys	Ser	Asp	Gly	Ala 50	Pro	Ser	Arg	Trp	Ser 55	Leu	Trp	Ser	Arg	Met 60
Arg	Arg	Trp	Gly	Cys 65	Pro	Leu	Arg	Leu	Ala 70	Leu	Ser	His	His	His 75
Leu	Arg	Pro	Arg	Thr 80	Val	Ser	Leu	Arg	Ser 85	Glu	Ala	Cys	Trp	Pro 90
Lys	Val	Cys	Gly	Leu 95	Arg	Ala	Pro	His	Gln 100	Pro	Ala	Pro	Суѕ	Ser 105
Thr	Gly	Pro	Pro	Leu 110	Gly	Arg	Val	Pro	Ser 115	Leu	Arg	Pro	Pro	Pro 120
Arg	Pro	Pro	Arg	Arg 125	Leu	Pro	His	Pro	Ser 130	Ser	Ile	Ser	Cys	Leu 135
Glu	Arg	Leu	Trp	Thr 140	Leu	Gly	Pro	Pro	Ser 145	Pro	Ala	Thr	Arg	Arg 150
Leu	Glu	Ser	Arg	Cys 155	Pro	Ala	Pro	Ala	Ala 160	Thr	Pro	Pro	Ser	Thr 165
Pro	Pro	Pro	Arg	Xaa 170	Xaa	Phe	Lys	Gly	Cys 175	Lys	Asn			

- (2) INFORMATION FOR SEQ ID NO: 40:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 197 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
      (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: LEUKNOT03
    - (B) CLONE: 1880830
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:

Met Ile Thr Cys Arg Val Cys Gln Ser Leu Ile Asn Val Glu Gly 10 15 Lys Met His Gln His Val Val Lys Cys Gly Val Cys Asn Glu Ala 25

Thr Pro Ile Lys Asn Ala Pro Pro Gly Lys Lys Tyr Val Arg Cys 40 35 Pro Cys Asn Cys Leu Leu Ile Cys Lys Val Thr Ser Gln Arg Ile 55 50 Ala Cys Pro Arg Pro Tyr Cys Lys Arg Ile Ile Asn Leu Gly Pro 70 65 Val His Pro Gly Pro Leu Ser Pro Glu Pro Gln Pro Met Gly Val 85 80 Arg Val Ile Cys Gly His Cys Lys Asn Thr Phe Leu Trp Thr Glu 95 100 Phe Thr Asp Arg Thr Leu Ala Arg Cys Pro His Cys Arg Lys Val 115 110 Ser Ser Ile Gly Arg Arg Tyr Pro Arg Lys Arg Cys Ile Cys Cys 130 135 125 Phe Leu Leu Gly Leu Leu Ala Val Thr Ala Thr Gly Leu Ala 145 140 Phe Gly Thr Trp Lys His Ala Arg Arg Tyr Gly Gly Ile Tyr Ala 165 160 155 Ala Trp Ala Phe Val Ile Leu Leu Ala Val Leu Cys Leu Gly Arg 175 170 Ala Leu Tyr Trp Ala Cys Met Lys Val Ser His Pro Val Gln Asn 185 Phe Ser

# (2) INFORMATION FOR SEQ ID NO: 41:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 302 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: OVARNOT07
- (B) CLONE: 1905325
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:

Met Leu Lys Asp Ile Ile Lys Glu Tyr Thr Asp Val Tyr Pro Glu 10 Ile Ile Glu Arg Ala Gly Tyr Ser Leu Glu Lys Val Phe Gly Ile 25 20 Gln Leu Lys Glu Ile Asp Lys Asn Asp His Leu Tyr Ile Leu Leu 40 35 Ser Thr Leu Glu Pro Thr Asp Ala Gly Ile Leu Gly Thr Thr Lys 55 50 Asp Ser Pro Lys Leu Gly Leu Leu Met Val Leu Leu Ser Ile Ile 65 70 Phe Met Asn Gly Asn Arg Ser Ser Glu Ala Val Ile Trp Glu Val 85 80 Leu Arg Lys Leu Gly Leu Arg Pro Gly Ile His His Ser Leu Phe 105 100 95 Gly Asp Val Lys Lys Leu Ile Thr Asp Glu Phe Val Lys Gln Lys 120 115 110 Tyr Leu Asp Tyr Ala Arg Val Pro Asn Ser Asn Pro Pro Glu Tyr 130 125 Glu Phe Phe Trp Gly Leu Arg Ser Tyr Tyr Glu Thr Ser Lys Met 145 140 Lys Val Leu Lys Phe Ala Cys Lys Val Gln Lys Lys Asp Pro Lys

				155					160					165
Glu	Trp	Ala	Ala	Gln 170	Tyr	Arg	Glu	Ala	Met 175	Glu	Ala	Asp	Leu	Lys 180
Ala	Ala	Ala	Glu	Ala 185	Ala	Ala	Glu	Ala	Lys 190	Ala	Arg	Ala	Glu	Ile 195
Arg	Ala	Arg	Met		Ile	Gly	Leu	Gly	Ser 205	Glu	Asn	Ala	Ala	Gly 210
Pro	Cys	Asn	Trp	Asp	Glu	Ala	Asp	Ile	Gly 220	Pro	Trp	Ala	Lys	Ala 225
Arg	Ile	Gln	Ala	Gly 230	Ala	Glu	Ala	Lys	Ala 235	Lys	Ala	Gln	Glu	Ser 240
Gly	Ser	Ala	Ser	Thr 245	Gly	Ala	Ser	Thr	Ser 250	Thr	Asn	Asn	Ser	Ala 255
Ser	Ala	Ser	Ala	Ser 260	Thr	Ser	Gly	Gly	Phe 265	Ser	Ala	Gly	Ala	Ser 270
Leu	Thr	Ala	Thr		Thr	Phe	Gly	Leu	Phe 280	Ala	Gly	Leu	Gly	Gly 285
Ala	Gly	Ala	Ser	Thr 290	Ser	Gly	Ser	Ser	Gly 295	Ala	Cys	Gly	Phe	Ser 300
Tyr	Lys													

# (2) INFORMATION FOR SEQ ID NO: 42:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 164 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRSTTUT01 (B) CLONE: 1919931

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:

Met	Arg	Thr	Leu	Glu 5	Asn	Gln	Gly	Phe	Lys 10	Ile	Leu	Pro	Phe	Leu 15
Gly	Val	Lys	Glu	Val 20	Trp	Gln	Lys	Gln	Asn 25	Lys	Leu	Ile	Ser	Arg 30
			Cys	35					40					45
			Val	50					55					60
Cys	Phe	Leu	Phe	His 65	Pro	Trp	Thr	Leu	Gln 70	Glu	Ile	Ala	Pro	Cys 75
Phe	Cys	Leu	Cys	Ile 80	Thr	Glu	Lys	Gly	Ser 85	Met	Lys	Val	Ala	Gln 90
Val	Arg	Pro	Phe	His 95	Cys	Pro	Pro	Gly	Ala 100	Gly	Phe	Ala	Leu	Pro 105
			Leu	110					115					Leu 120
			Gln	125					130					135
Ser	Thr	Cys	His	Cys 140	Val	Cys	Lys	Ser	Ser 145	Phe	Ser	Phe	Phe	Leu 150
Ala	His	Leu	Thr	Leu 155	Val	Met	Ser	Leu	Ile 160	Thr	Thr	Thr	Ile	

				155					160					165
Glu	Trp	Ala	Ala	Gln 170	Tyr	Arg	Glu	Ala	Met 175	Glu	Ala	Asp	Leu	Lys 180
Ala	Ala	Ala	Glu		Ala	Ala	Glu	Ala		Ala	Arg	Ala	Glu	Ile 195
Arg	Ala	Arg	Met		Ile	Gly	Leu	Gly	Ser 205	Glu	Asn	Ala	Ala	Gly 210
Pro	Cys	Asn	Trp		Glu	Ala	Asp	Ile	Gly 220	Pro	Trp	Ala	Lys	Ala 225
Arg	Ile	Gln	Ala	Gly 230	Ala	Glu	Ala	Lys	Ala 235	Lys	Ala	Gln	Glu	Ser 240
Gly	Ser	Ala	Ser	Thr 245	Gly	Ala	Ser	Thr	Ser 250	Thr	Asn	Asn	Ser	Ala 255
Ser	Ala	Ser	Ala	Ser 260	Thr	Ser	Gly	Gly	Phe 265	Ser	Ala	Gly	Ala	Ser 270
Leu	Thr	Ala	Thr		Thr	Phe	Gly	Leu	Phe 280	Ala	Gly	Leu	Gly	Gly 285
Ala	Gly	Ala	Ser	Thr 290	Ser	Gly	Ser	Ser	Gly 295	Ala	Cys	Gly	Phe	Ser 300
Tyr	Lys													

# (2) INFORMATION FOR SEQ ID NO: 42:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 164 amino acids

  - (B) TYPE: amino acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRSTTUT01
- (B) CLONE: 1919931

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:

Met	Arg	Thr	Leu	Glu 5	Asn	Gln	Gly	Phe	Lys 10	Ile	Leu	Pro	Phe	Leu 15
_	Val			20					25					30
	Ile			35					40					45
	Ile			50					55					60
_	Phe			65					70					75
	Cys			80					85					Gln 90
Val	Arg	Pro	Phe	His 95	Cys	Pro	Pro	Gly	Ala 100	Gly	Phe	Ala	Leu	Pro 105
	Leu			110					115					Leu 120
His	Ile	Ser	Gln	Val 125	Ser	Ala	Gln	Lys	Ser 130	Pro	Phe	Gly	Gly	Val 135
Ser	Thr	Cys	His	Cys 140	Val	Cys	Lys	Ser	Ser 145	Phe	Ser	Phe	Phe	Leu 150
Ala	His	Leu	Thr	Leu 155	Val	Met	Ser	Leu	Ile 160	Thr	Thr	Thr	Ile	

# (2) INFORMATION FOR SEQ ID NO: 43

- (i) SEQUENCE CHARACTERISTICS:
  - $(\bar{A})$  LENGTH: 235 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRSTNOT04
- (B) CLONE: 1969426
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:

```
Met Ser Pro Thr Leu Ser Ser Ile Thr Gln Gly Val Pro Leu Asp
                                      10
                  5
Thr Ser Lys Leu Ser Thr Asp Gln Arg Leu Pro Pro Tyr Pro Tyr
                                      25
                 2.0
Ser Ser Pro Ser Leu Val Leu Pro Thr Gln Pro His Thr Pro Lys
                 35
                                      40
Ser Leu Gln Gln Pro Gly Leu Pro Ser Gln Ser Cys Ser Val Gln
                                      55
                 50
Ser Ser Gly Gly Gln Pro Pro Gly Arg Gln Ser His Tyr Gly Thr
                                      70
                 65
Pro Tyr Pro Pro Gly Pro Ser Gly His Gly Gln Gln Ser Tyr His
                                      85
                 80
Arg Pro Met Ser Asp Phe Asn Leu Gly Asn Leu Glu Gln Phe Ser
                                     100
                 95
Met Glu Ser Pro Ser Ala Ser Leu Val Leu Asp Pro Pro Gly Phe
                                                          120
                                     115
                110
Ser Glu Gly Pro Gly Phe Leu Gly Gly Glu Gly Pro Met Gly Gly
                                                          135
                                     130
                 125
Pro Gln Asp Pro His Thr Phe Asn His Gln Asn Leu Thr His Cys
                                                          150
                                     145
                140
Ser Arg His Gly Ser Gly Pro Asn Ile Ile Leu Thr Gly Asp Ser
                                                          165
                                     160
                 155
Ser Pro Gly Phe Ser Lys Glu Ile Ala Ala Ala Leu Ala Gly Val
                                                          180
                                     175
                 170
Pro Gly Phe Glu Val Ser Ala Ala Gly Leu Glu Leu Gly Leu Gly
                                                          195
                                     190
                 185
Leu Glu Asp Glu Leu Arg Met Glu Pro Leu Gly Leu Glu Gly Leu
                 200
                                     205
Asn Met Leu Ser Asp Pro Cys Ala Leu Leu Pro Asp Pro Ala Val
                                     220
                 215
Glu Glu Ser Phe Arg Ser Asp Arg Leu Gln
                 230
```

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 203 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: UCMCL5T01



(B) CLONE: 1969948

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 44:

Met Asn Tyr Phe Pro Leu Ala Pro Phe Asn Gln Leu Leu Gln Lys Asp Ile Ile Ser Glu Leu Leu Thr Ser Asp Asp Met Lys Asn Ala 25 20 Tyr Lys Leu His Thr Leu Asp Thr Cys Leu Lys Leu Asp Asp Thr 40 35 Val Tyr Leu Arg Asp Ile Ala Leu Ser Leu Pro Gln Leu Pro Arg 55 50 Glu Leu Pro Ser Ser His Thr Asn Ala Lys Val Ala Glu Val Leu 70 65 Ser Ser Leu Leu Gly Gly Glu Gly His Phe Ser Lys Asp Val His 80 85 Leu Pro His Asn Tyr His Ile Asp Phe Glu Ile Arg Met Asp Thr 100 95 Asn Arg Asn Gln Val Leu Pro Leu Ser Asp Val Asp Thr Thr Ser 120 115 110 Ala Thr Asp Ile Gln Arg Val Ala Val Leu Cys Val Ser Arg Ser 130 125 Ala Tyr Cys Leu Gly Ser Ser His Pro Arg Gly Phe Leu Ala Met 150 140 145 Lys Met Arg His Leu Asn Ala Met Gly Phe His Val Ile Leu Val 160 155 Asn Asn Trp Glu Met Asp Lys Leu Glu Met Glu Asp Ala Val Thr 175 170 Phe Leu Lys Thr Lys Ile Tyr Ser Val Glu Ala Leu Pro Val Ala 185 Ala Val Asn Val Gln Ser Thr Gln 200

### (2) INFORMATION FOR SEQ ID NO: 45:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 359 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LUNGAST01
  - (B) CLONE: 1988911
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 45:

Met Glu Arg Gly Asn Val Leu Ser Arg Ala Pro Ser Arg Ala His 10 Gly Thr His Phe Gly Asp Asp Arg Phe Glu Asp Leu Glu Glu Ala 30 25 20 Asn Pro Phe Ser Phe Arg Glu Phe Leu Lys Thr Lys Asn Leu Gly 45 35 40 Leu Ser Lys Glu Asp Pro Ala Ser Arg Ile Tyr Ala Lys Glu Ala 55 50 Ser Arg His Ser Leu Gly Leu Asp His Asn Ser Pro Pro Ser Gln 70 Thr Gly Gly Tyr Gly Leu Glu Tyr Gln Gln Pro Phe Phe Glu Asp

				80					85					90
Pro	Thr	Gly	Ala	Gly 95	Asp	Leu	Leu	Asp	Glu 100	Glu	Glu	Asp	Glu	Asp 105
Thr	Gly	Trp	Ser	Gly 110	Ala	Tyr	Leu	Pro	Ser 115	Ala	Ile	Glu	Gln	Thr 120
His	Pro	Glu	Arg	Val 125	Pro	Ala	Gly	Thr	Ser 130	Pro	Cys	Ser	Thr	Tyr 135
Leu	Ser	Phe	Phe	Ser 140	Thr	Pro	Ser	Glu	Leu 145	Ala	Gly	Pro	Glu	Ser 150
Leu	Pro	Ser	Trp	Ala 155	Leu	Ser	Asp	Thr	Asp 160	Ser	Arg	Val	Ser	Pro 165
Ala	Ser	Pro	Ala	Gly 170	Ser	Pro	Ser	Ala	Asp 175	Phe	Ala	Val	His	Gly 180
Glu	Ser	Leu	Gly	Asp 185	Arg	His	Leu	Arg	Thr 190	Leu	Gln	Ile	Ser	Tyr 195
Asp	Ala	Leu	Lys	Asp 200	Glu	Asn	Ser	Lys	Leu 205	Arg	Arg	Lys	Leu	Asn 210
			Ser	215					220					225
Leu	Glu	Arg	Lys	Leu 230	Glu	Ala	Lys	Met	Ile 235	Lys	Glu	Glu	Ser	Asp 240
_			Leu	245					250					Leu 255
			Thr	260					265					Val 270
Lys	Leu	Lys	Gln	Glu 275	Ile	Ser	Leu	Leu	Gln 280	Ala	Gln	Val	Ser	Asn 285
		_	Glu	290					295					300
Leu	Thr	Val	Val	Lys 305	Gln	Asn	Ala	Asp	Val 310					315
_			Met	320					325		_		Leu	330
			Glu	335					340					Ser 345
Ile	Asp	Arg	Ile	Ser 350	Glu	Val	Lys	Asp	Glu 355	Glu	Glu	Asp	Ser	

### (2) INFORMATION FOR SEQ ID NO: 46:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 150 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: OVARNOT03
- (B) CLONE: 2061561

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 46:

Met Gly Gly Lys Pro His Lys Glu Pro Arg Ala Lys Gly Pro Leu 10 Ser Ile Phe Tyr Pro Gly Ser Thr Ala Pro Val Ile Thr Gln Arg 30 20 25 Thr Pro Xaa Ala Ala Leu Lys Pro Pro Pro Ile Lys Gly Ala Gly 35 40

Pro Thr Ile Ala Pro Ile Lys Gly Xaa Xaa Asn Phe Gly Lys Arg 55 Pro Thr Val Thr Xaa Pro Xaa Trp Xaa Ile Ser Pro Asn Trp Gly 75 70 65 Lys Arg Gly Xaa Cys Xaa Xaa Xaa Gly Ile Lys Trp Val Xaa Pro 80 85 Arg Val Ser Gln Ala Arg Thr Phe Lys Thr Thr Ala Asn Glu Leu 100 95 Xaa Phe Xaa Asp Thr Phe Glu Glu Xaa Xaa Arg Xaa Xaa His Ala 110 115 Xaa Val Ser Xaa Glu Pro Gln Pro Arg Cys Pro Leu Gly Glu Ser 130 125 Arg Ser Leu Gly Ala Ala Val Cys Arg Trp Asp Ser Phe Asp Phe 145 140

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 402 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: PANCNOT04
- (B) CLONE: 2084489
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 47:

Met	Pro	Pro	Val	Ser 5	Arg	Ser	Ser	Tyr	Ser 10	Glu	Asp	Ile	Val	Gly 15
Ser	Arg	Arg	Arg	Arg 20	Arg	Ser	Ser	Ser	Gly 25	Ser	Pro	Pro	Ser	Pro 30
Gln	Ser	Arg	Cys	Ser 35	Ser	Trp	Asp	Gly	Cys 40	Ser	Arg	Ser	His	Ser 45
Arg	Gly	Arg	Glu	Gly 50	Leu	Arg	Pro	Pro	Trp 55	Ser	Glu	Leu	Asp	Val 60
Gly	Ala	Leu	Tyr	Pro 65	Phe	Ser	Arg	Ser	Gly 70	Ser	Arg	Gly	Arg	Leu 75
Pro	Arg	Phe	Arg	Asn 80	Tyr	Ala	Phe	Ala	Ser 85	Ser	Trp	Ser	Thr	Ser 90
Tyr	Ser	Gly	Tyr	Arg 95	Tyr	His	Arg	His	Cys 100	Tyr	Ala	Glu	Glu	Arg 105
Gln	Ser	Ala	Glu	Asp 110	Tyr	Glu	Lys	Glu	Glu 115	Ser	His	Arg	Gln	Arg 120
Arg	Leu	Lys	Glu	Arg 125	Glu	Arg	Ile	Gly	Glu 130	Leu	Gly	Ala	Pro	Glu 135
Val	Trp	Gly	Pro	Ser 140	Pro	Lys	Phe	Pro	Gln 145	Leu	Asp	Ser	Asp	Glu 150
His	Thr	Pro	Val	Glu 155	Asp	Glu	Glu	Glu	Val 160	Thr	His	Gln	Lys	Ser 165
Ser	Ser	Ser	Asp	Ser 170	Asn	Ser	Glu	Glu	His 175	Arg	Lys	Lys	Lys	Thr 180
Ser	Arg	Ser	Arg	Asn 185	Lys	Lys	Lys	Arg	Lys 190	Asn	Lys	Ser	Ser	Lys 195
Arg	Lys	His	Arg	Lys	Tyr	Ser	Asp	Ser	Asp	Ser	Asn	Ser	Glu	Ser

				200					205					210
Asp	Thr	Asn	Ser	Asp 215	Ser	Asp	Asp	Asp	Lys 220	Lys	Arg	Val	Lys	Ala 225
Lys	Lys	Lys	Lys	Lys 230	Lys	Lys	Lys	His	Lys 235	Thr	Lys	Lys	Lys	Lys 240
	_			245				Ser	250					Asp 255
Ser	Glu	Glu	Asp	Leu 260	Ser	Glu	Ala	Thr	Trp 265	Met	Glu	Gln	Pro	Asn 270
				275				Gly	280					Ile 285
				290				Leu	295					300
				305				Ala	310					315
				320				Ile	325					330
Ile	Gly	Ser	Phe	Glu 335	Cys	Ser	Gly	Tyr	Val 340	Met	Ser	Gly	Ser	Arg 345
	_	_		350				Leu	355					360
_				365				Leu	370					375
Glu	Arg	Arg	Lys	Arg 380	Glu	Ser	Lys	Ile	Leu 385	Ala	Ser	Phe	Arg	Glu 390
Met	Val	His	Lys	Lys 395	Thr	Lys	Glu	Lys	Asp 400	Asp	Lys			

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:

  (A) LENGTH: 311 amino acids

  (B) TYPE: amino acid

  (C) STRANDEDNESS: single

  (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: SPLNFET02 (B) CLONE: 2203226

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 48:

	His			5					10					15
Leu	Pro	Ser	Lys	Arg 20	Arg	Ile	Pro	Val	Ser 25	Gln	Pro	Gly	Met	Ala 30
Asp	Pro	His	Gln	Leu 35	Phe	Asp	Asp	Thr	Ser 40	Ser	Ala	Gln	Ser	Arg 45
Gly	Tyr	Gly	Ala	Gln 50	Arg	Ala	Pro	Gly	Gly 55	Leu	Ser	Tyr	Pro	Ala 60
Ala	Ser	Pro	Thr	Pro 65	His	Ala	Ala	Phe	Leu 70	Ala	Asp	Pro	Val	Ser 75
Asn	Met	Ala	Met	Ala 80	Tyr	Gly	Ser	Ser	Leu 85	Ala	Ala	Gln	Gly	Lys 90
Glu	Leu	Val	Asp	Lys 95	Asn	Ile	Asp	Arg	Phe 100	Ile	Pro	Ile	Thr	Lys 105
Leu	Lys	Tyr	Tyr	Phe 110	Ala	Val	Asp	Thr	Met 115	Tyr	Val	Gly	Arg	Lys 120

48:

Leu Gly Leu Leu Phe Phe Pro Tyr Leu His Gln Asp Trp Glu Val 130 125 Gln Tyr Gln Gln Asp Thr Pro Val Ala Pro Arg Phe Asp Val Asn 145 140 Ala Pro Asp Leu Tyr Ile Pro Ala Met Ala Phe Ile Thr Tyr Val 160 155 Leu Val Ala Gly Leu Ala Leu Gly Thr Gln Asp Arg Phe Ser Pro 175 170 Asp Leu Leu Gly Leu Gln Ala Ser Ser Ala Leu Ala Trp Leu Thr 190 185 Leu Glu Val Leu Ala Ile Leu Leu Ser Leu Țyr Leu Val Thr Val 205 Asn Thr Asp Leu Thr Thr Ile Asp Leu Val Ala Phe Leu Gly Tyr 220 215 Lys Tyr Val Gly Met Ile Gly Gly Val Leu Met Gly Leu Leu Phe 235 230 Gly Lys Ile Gly Tyr Tyr Leu Val Leu Gly Trp Cys Cys Val Ala 250 245 Ile Phe Val Phe Met Ile Arg Thr Leu Arg Leu Lys Ile Leu Ala 265 260 Asp Ala Ala Glu Gly Val Pro Val Arg Gly Ala Arg Asn Gln 280 275 Leu Arg Met Tyr Leu Thr Met Ala Val Ala Ala Ala Gln Pro Met 290 295 Leu Met Tyr Trp Leu Thr Phe His Leu Val Arg 305

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 316 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: PROSNOT16
  - (B) CLONE: 2232884
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 49:

Met Ala Ser Ala Asp Glu Leu Thr Phe His Glu Phe Glu Glu Ala Thr Asn Leu Leu Ala Asp Thr Pro Asp Ala Ala Thr Thr Ser Arg 25 Ser Asp Gln Leu Thr Pro Gln Gly His Val Ala Val Ala Val Gly 40 35 Ser Gly Gly Ser Tyr Gly Ala Glu Asp Glu Val Glu Glu Ger 50 55 Asp Lys Ala Ala Leu Leu Gln Glu Gln Gln Gln Gln Gln Pro 70 65 Gly Phe Trp Thr Phe Ser Tyr Tyr Gln Ser Phe Phe Asp Val Asp 90 80 8.5 Thr Ser Gln Val Leu Asp Arg Ile Lys Gly Ser Leu Leu Pro Arg 100 95 Pro Gly His Asn Phe Val Arg His His Leu Arg Asn Arg Pro Asp 110 115 Leu Tyr Gly Pro Phe Trp Ile Cys Ala Thr Leu Ala Phe Val Leu

				125					130					135
				140	Leu				145					Asp 150
Pro	Ser	Ile	His	Tyr 155	Ser	Pro	Gln	Phe	His 160	Lys	Val	Thr	Val	Ala 165
_				170	Cys				175					180
				185	Trp				190					Gly 195
	_			200	Glu				205					Leu 210
Phe	Val	Phe	Ile	Pro 215	Met	Val	Val	Leu	Trp 220	Leu	Ile	Pro	Val	Pro 225
_				230	Phe				235					240
Ala	Gly	Leu	Val	Phe 245	Thr	Leu	Trp	Pro	Val 250	Val	Arg	Glu	Asp	Thr 255
Arg	Leu	Val	Ala	Thr 260	Val	Leu	Leu	Ser	Val 265	Val	Val	Leu	Leu	His 270
				275	Gly				280					285
Pro	Pro	Glu	Asn	Val 290	Ala	Pro	Pro	Pro	Gln 295	Ile	Thr	Ser	Leu	Pro 300
Ser	Asn	Ile	Ala	Leu 305	Ser	Pro	Thr	Leu	Pro 310	Gln	Ser	Leu	Ala	Pro 315
Ser														

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 346 amino acids

  - (B) TYPE: amino acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: COLNNOT11 (B) CLONE: 2328134
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 50:

Met	Thr	Pro	Arg	Thr 5	Trp	Trp	Pro	Arg	Pro 10	Ala	Gly	Trp	Gly	Thr 15
Cys	Arg	Ala	Ala	Gly 20	Trp	Pro	Arg	Ser	Val 25	Pro	Trp	Ala	Arg	Thr 30
Ala	Ala	Ser	Leu	Val 35	Phe	Val	Pro	Thr	Arg 40	Arg	Arg	Ser	Gly	Pro 45
Ser	Gly	Thr	Ala	Ser 50	Val	Ala	Ala	Met	Ala 55	Tyr	His	Ser	Gly	Tyr 60
Gly	Ala	His	Gly	Ser 65	Lys	His	Arg	Ala	Arg 70	Ala	Ala	Pro	Asp	Pro 75
Pro	Pro	Leu	Phe	Asp 80	Asp	Thr	Ser	Gly	Gly 85	Tyr	Ser	Ser	Gln	Pro 90
Gly	Gly	Tyr	Pro	Ala 95	Thr	Gly	Ala	Asp	Val 100	Ala	Phe	Ser	Val	Asn 105
His	Leu	Leu	Gly	Asp 110	Pro	Met	Ala	Asn	Val 115	Ala	Met	Ala	Tyr	Gly 120
Ser	Ser	Ile	Ala	Ser 125	His	Gly	Lys	Asp	Met 130	Val	His	Lys	Glu	Leu 135

```
His Arg Phe Val Ser Val Ser Lys Leu Lys Tyr Phe Phe Ala Val
                                    145
                140
Asp Thr Ala Tyr Val Ala Lys Lys Leu Gly Leu Leu Val Phe Pro
                                    160
                155
Tyr Thr His Gln Asn Trp Glu Val Gln Tyr Ser Arg Asp Ala Pro
                                                         180
                                     175
                170
Leu Pro Pro Arg Gln Asp Leu Asn Ala Pro Asp Leu Tyr Ile Pro
                                    190
                185
Thr Met Ala Phe Ile Thr Tyr Val Leu Leu Ala Gly Met Ala Leu
                                     205
                200
Gly Ile Gln Lys Arg Phe Ser Pro Glu Val Leu Gly Leu Cys Ala
                                     220
                215
Ser Thr Ala Leu Val Trp Val Val Met Glu Val Leu Ala Leu Leu
                                     235
                230
Leu Gly Leu Tyr Leu Ala Thr Val Arg Ser Asp Leu Ser Thr Phe
                                                         255
                                     250
                245
His Leu Leu Ala Tyr Ser Gly Tyr Lys Tyr Val Gly Met Ile Leu
                                                          270
                260
                                     265
Ser Val Leu Thr Gly Leu Leu Phe Gly Ser Asp Gly Tyr Tyr Val
                                     280
                                                          285
                275
Ala Leu Ala Trp Thr Ser Ser Ala Leu Met Tyr Phe Ile Val Arg
                                                          300
                290
                                     295
Ser Leu Arg Thr Ala Ala Leu Gly Pro Asp Ser Met Gly Gly Pro
                                                          315
                305
                                     310
Val Pro Arg Gln Arg Leu Gln Leu Tyr Leu Thr Leu Gly Ala Ala
                                     325
                                                          330
                320
Ala Phe Gln Pro Leu Ile Ile Tyr Trp Leu Thr Phe His Leu Val
                                     340
                335
Arg
```

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 299 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: ISLTNOT01
  - (B) CLONE: 2382718
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 51:

Met Gly Thr Lys Ala Gln Val Glu Arg Lys Leu Cys Leu Phe 10 5 Ile Leu Ala Ile Leu Leu Cys Ser Leu Ala Leu Gly Ser Val Thr 25 -30 2.0 Val His Ser Ser Glu Pro Glu Val Arg Ile Pro Glu Asn Asn Pro 35 40 Val Lys Leu Ser Cys Ala Tyr Ser Gly Phe Ser Ser Pro Arg Val 60 50 55 Glu Trp Lys Phe Asp Gln Gly Asp Thr Thr Arg Leu Val Cys Tyr 7.0 Asn Asn Lys Ile Thr Ala Ser Tyr Glu Asp Arg Val Thr Phe Leu 85 90 -80 Pro Thr Gly Ile Thr Phe Lys Ser Val Thr Arg Glu Asp Thr Gly 95 100 Thr Tyr Thr Cys Met Val Ser Glu Glu Gly Gly Asn Ser Tyr Gly

51:

				110					115					120
Glu	Val	Lys	Val	Lys 125	Leu	Ile	Val	Leu	Val 130	Pro	Pro	Ser	Lys	Pro 135
Thr	Val	Asn	Ile	Pro 140	Ser	Ser	Ala	Thr	Ile 145	Gly	Asn	Arg	Ala	Val 150
Leu	Thr	Cys	Ser	Glu 155	Gln	Asp	Gly	Ser	Pro 160	Pro	Ser	Glu	Tyr	Thr 165
Trp	Phe	Lys	Asp	Gly 170	Ile	Val	Met	Pro	Thr 175	Asn	Pro	Lys	Ser	Thr 180
Arg	Ala	Phe	Ser	Asn 185	Ser	Ser	Tyr	Val	Leu 190	Asn	Pro	Thr	Thr	Gly 195
Glu	Leu	Val	Phe	Asp 200	Pro	Leu	Ser	Ala	Ser 205	Asp	Thr	Gly	Glu	Tyr 210
Ser	Cys	Glu	Ala	Arg 215	Asn	Gly	Tyr	Gly	Thr 220	Pro	Met	Thr	Ser	Asn 225
Ala	Val	Arg	Met	Glu 230	Ala	Val	Glu	Arg	Asn 235	Val	Gly	Val	Ile	Val 240
				245					250				Val	255
_				260					265				Thr	270
Lys	Gly	Thr	Ser	Ser 275	Lys	Lys	Val	Ile	Tyr 280	Ser	Gln	Pro	Ser	Ala 285
Arg	Ser	Glu	Gly	Glu 290	Phe	Lys	Gln	Thr	Ser 295	Ser	Phe	Leu	Val	

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - $(\bar{A})$  LENGTH: 351 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
    (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: ENDANOT01
- (B) CLONE: 2452208

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 52:

Met Ala	Ser	Thr	Gly 5	Ser	Gln	Ala	Ser	Asp 10	Ile	Asp	Glu	Ile	Phe 15
Gly Phe	Phe	Asn	Asp 20	Gly	Glu	Pro	Pro	Thr 25	Lys	Lys	Pro	Arg	Lys 30
Leu Leu	Pro	Ser	Leu 35	Lys	Thr	Lys	Lys	Pro 40	Arg	Glu	Leu	Val	Leu 45
Val Ile	Gly	Thr	Gly 50	Ile	Ser	Ala	Ala	Val 55	Ala	Pro	Gln	Val	Pro 60
Ala Leu	Lys	Ser	Trp 65	Lys	Gly	Leu	Ile	Gln 70	Ala	Leu	Leu	Asp	Ala 75
Ala Ile	Asp	Phe	Asp 80	Leu	Leu	Glu	Asp	Glu 85	Glu	Ser	Lys	Lys	Phe 90
Gln Lys	Cys	Leu	His 95	Glu	Asp	Lys	Asn	Leu 100	Val	His	Val	Ala	His 105
Asp Leu	Ile	Gln	Lys 110	Leu	Ser	Pro	Arg	Thr 115	Ser	Asn	Val	Arg	Ser 120
Thr Phe	Phe	Lys	Asp 125	Cys	Leu	Tyr	Glu	Val 130	Phe	Asp	Asp	Leu	Glu 135

Ser Lys Met Glu Asp Ser Gly Lys Gln Leu Leu Gln Ser Val Leu 145 140 His Leu Met Glu Asn Gly Ala Leu Val Leu Thr Thr Asn Phe Asp 160 155 Asn Leu Leu Glu Leu Tyr Ala Ala Asp Gln Gly Lys Gln Leu Glu 175 170 Ser Leu Asp Leu Thr Asp Glu Lys Lys Val Leu Glu Trp Ala Gln 195 190 185 Glu Lys Arg Lys Leu Ser Val Leu His Ile His Gly Val Tyr Thr 205 210 200 Asn Pro Ser Gly Ile Val Leu His Pro Ala Gly Tyr Gln Asn Val 220 215 Leu Arg Asn Thr Glu Val Met Arg Glu Ile Gln Lys Leu Tyr Glu 235 230 Asn Lys Ser Phe Leu Phe Leu Gly Cys Gly Trp Thr Val Asp Asp 245 250 Thr Thr Phe Gln Ala Leu Phe Leu Glu Ala Val Lys His Lys Ser 265 260 Asp Leu Glu His Phe Met Leu Val Arg Arg Gly Asp Val Asp Glu 280 285 275 Phe Lys Lys Leu Arg Glu Asn Met Leu Asp Lys Gly Ile Lys Val 295 290 Ile Ser Tyr Gly Asp Asp Tyr Ala Asp Leu Pro Glu Tyr Phe Lys 305 310 315 Arg Leu Thr Cys Glu Ile Ser Thr Arg Gly Thr Ser Ala Gly Met 325 320 Val Arg Glu Gly Gln Leu Asn Gly Ser Ser Ala Ala His Ser Glu 345 340 335 Ile Arg Gly Cys Ser Thr 350

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 662 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: ENDANOT01
- (B) CLONE: 2457825

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 53:

Met Thr Ala Lys Lys Gln Cys Leu Leu Arg Leu Gly Val Leu Arg 1.0 Gln Asp Trp Pro Asp Thr Asn Arg Leu Leu Gly Ser Ala Asn Val 20 25 Val Pro Glu Ala Leu Gln Arg Phe Thr Arg Ala Ala Ala Asp Phe 40 35 Ala Thr His Gly Lys Leu Gly Lys Leu Glu Phe Ala Gln Asp Ala 50 55 His Gly Gln Pro Asp Val Ser Ala Phe Asp Phe Thr Ser Met Met 75 70 65 Arg Ala Glu Ser Ser Ala Arg Val Gln Glu Lys His Gly Ala Arg 85 80 Leu Leu Gly Leu Val Gly Asp Cys Leu Val Glu Pro Phe Trp

				95					100					105
		_		110					115			Ala		120
				125					130			Glu		135
				140					145			Leu		150
				155					160			Tyr		165
				170					175			Ala		TRO
				185					190			Lys		195
				200					205			Pro		210
_				215					220			Cys		225
				230					235			Leu		240
				245					250			Tyr		255
				260					265			Leu		2/0
				275					280			Asn		285
_				290					295			Ala		300
				305					310			His		315
				320					325			Gln		330
				335					340			Leu		345
				350					355			Asp		360
				365					370			Pro		3/5
				380					385	,		Thr		390
				395					400	}		Ala		405
				410					415	5		Asn		420
				425	,				430	)		Glu		435
				440	)				445	)		Arg		450
				455	·				460	)		s Glu		465
				470	)				475	5		o Val		480
_				485	5				490	)		a Lys		495
				500	)				505	5		Let		510
				515	5				520	)		s Ala		525
				530	)				53	5		a Aro		540
Glı	ı Ala	a Gli	ı GI	y Va. 545		s Lev	ı Gill	л ге	550 550		ı Arç	g Arq	y GII.	555 555

Ser Ser Pro Glu Gln Gln Lys Lys Leu Trp Val Gly Gln Leu Leu 560 565 Gln Leu Val Asp Lys Lys Asn Ser Leu Val Ala Glu Glu Ala Glu 580 575 Leu Met Ile Thr Val Gln Glu Leu Asn Leu Glu Glu Lys Gln Trp 590 595 Gln Leu Asp Gln Glu Leu Arq Gly Tyr Met Asn Arg Glu Glu Asn 610 605 Leu Lys Thr Ala Ala Asp Arg Gln Ala Glu Asp Gln Val Leu Arg 625 630 620 Lys Leu Val Asp Leu Val Asn Gln Arg Asp Ala Leu Ile Arg Phe 640 635 Gln Glu Glu Arg Arg Leu Ser Glu Leu Ala Leu Gly Thr Gly Ala 655 650 Gln Gly

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 115 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: THP1NOT03
- (B) CLONE: 2470740
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 54:

Met Ala Ser Trp Pro Ala Ser Pro Leu Gln Trp Gly Pro Pro Leu 10 Ala Ser Cys Pro Ser Cys Cys Cys Cys Cys Phe His Cys Trp Gln 30 20 25 Pro Arg Val Gly Val Ala Cys Arg Gln Arg Cys Trp Pro Leu Arg 40 35 Trp Gly Trp Trp Val Trp Gly Pro Pro Thr Cys Ser Phe Val Gln 50 5.5 Pro Cys Thr Cys Pro Pro Val Phe Ser Tyr Ser Trp Pro Arg Val 70 65 Pro His Trp Gly Pro Ser Trp Xaa Met Ser Trp Arg Arg Leu 80 85 Met Gly Val Pro Leu Gly Leu Trp Asn Cys Leu Val Leu Lys Leu 95 100 Xaa Gln Gly Leu Ala Pro Thr Ser Gly Gly 110

# (2) INFORMATION FOR SEQ ID NO: 55:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 157 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:

(A) LIBRARY: SMCANOT01
(B) CLONE: 2479092

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 55:

Met Glu Ala Leu Arg Arg Ala His Glu Val Ala Leu Arg Leu Leu 1.0 Leu Cys Arg Pro Trp Ala Ser Arg Ala Ala Ala Arg Pro Lys Pro 25 20 Ser Ala Ser Glu Val Leu Thr Arg His Leu Leu Gln Arg Arg Leu 40 Pro His Trp Thr Ser Phe Cys Val Pro Tyr Ser Ala Val Arg Asn 55 50 Asp Gln Phe Gly Leu Ser His Phe Asn Trp Pro Val Gln Gly Ala 65 70 Asn Tyr His Val Leu Arg Thr Gly Cys Phe Pro Phe Ile Lys Tyr 85 80 His Cys Ser Lys Ala Pro Trp Gln Asp Leu Ala Arg Gln Asn Arg 100 105 95 Phe Phe Thr Ala Leu Lys Val Val Asn Leu Gly Ile Pro Thr Leu 110 115 Leu Tyr Gly Leu Gly Ser Trp Leu Phe Ala Arg Val Thr Glu Thr 130 125 Val His Thr Ser Tyr Gly Pro Ile Thr Val Tyr Phe Leu Asn Lys 145 140 Glu Asp Glu Gly Ala Met Tyr 155

### (2) INFORMATION FOR SEQ ID NO: 5

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 197 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: SMCANOT01
  - (B) CLONE: 2480544
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 56:

Met Pro Pro Ala Gly Leu Arg Arg Ala Ala Pro Leu Thr Ala Ile 10 Ala Leu Leu Val Leu Gly Ala Pro Leu Val Leu Ala Gly Glu Asp 25 20 Cys Leu Trp Tyr Leu Asp Arg Asn Gly Ser Trp His Pro Gly Phe 40 45 35 Asn Cys Glu Phe Phe Thr Phe Cys Cys Gly Thr Cys Tyr His Arg 55 50 Tyr Cys Cys Arg Asp Leu Thr Leu Leu Ile Thr Glu Arg Gln Gln 70 65 Lys His Cys Leu Ala Phe Ser Pro Lys Thr Ile Ala Gly Ile Ala 90 80 -85 Ser Ala Val Ile Leu Phe Val Ala Val Val Ala Thr Thr Ile Cys 100 105 95 Cys Phe Leu Cys Ser Cys Cys Tyr Leu Tyr Arg Arg Gln Gln 115 110

```
Leu Gln Ser Pro Phe Glu Gly Gln Glu Ile Pro Met Thr Gly Ile
                125
                                    130
Pro Val Gln Pro Val Tyr Pro Tyr Pro Gln Asp Pro Lys Ala Gly
                                    145
                140
Pro Ala Pro Pro Gln Pro Gly Phe Met Tyr Pro Pro Ser Gly Pro
                155
                                    160
Ala Pro Gln Tyr Pro Leu Tyr Pro Ala Gly Pro Pro Val Tyr Asn
                                    175
                170
Pro Ala Ala Pro Pro Pro Tyr Met Pro Pro Gln Pro Ser Tyr Pro
                                    190
                                                        195
Gly Ala
```

### (2) INFORMATION FOR SEQ ID NO: 57:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 245 amino acids

  - (B) TYPE: amino acid(C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: BRAITUT21
  - (B) CLONE: 2518547
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 57:

Met	Gly	Gly	Ala	Ser 5	Arg	Arg	Val	Glu	Ser 10	Gly	Ala	Trp	Ala	Tyr 15
Leu	Ser	Pro	Leu	Val 20	Leu	Arg	Lys	Glu	Leu 25	Glu	Ser	Leu	Val	Glu 30
Asn	Glu	Gly	Ser	Glu 35	Val	Leu	Ala	Leu	Pro 40	Glu	Leu	Pro	Ser	Ala 45
His	Pro	Ile	Ile	Phe 50	Trp	Asn	Leu	Leu	Trp 55	Tyr	Phe	Gln	Arg	Leu 60
Arg	Leu	Pro	Ser	Ile 65	Leu	Pro	Gly	Leu	Val 70	Leu	Ala	Ser	Cys	Asp 75
Gly	Pro	Ser	His	Ser 80	Gln	Ala	Pro	Ser	Pro 85	Trp	Leu	Thr	Pro	Asp 90
Pro	Ala	Ser	Val	Gln 95	Val	Arg	Leu	Leu	Trp 100	Asp	Val	Leu	Thr	Pro 105
-			Ser	110					115					His 120
			Pro	125					130					Ala 135
Ser	Leu	Ser	Leu	Ala 140	Leu	Leu	Glu	Ser	Val 145	Leu	Arg	His	Val	Gly 150
Leu	Asn	Glu	Val	His 155	Lys	Ala	Val	Gly	Leu 160	Leu	Leu	Glu	Thr	Leu 165
Gly	Pro	Pro	Pro	Thr 170	Gly	Leu	His	Leu	Gln 175	Arg	Gly	Ile	Tyr	Arg 180
Glu	Ile	Leu	Phe	Leu 185	Thr	Met	Ala	Ala	Leu 190	Gly	Lys	Asp	His	Val 195
Asp	Ile	Val	Ala	Phe 200	Asp	Lys	Lys	Tyr	Lys 205	Ser	Ala	Phe	Asn	Lys 210
Leu	Ala	Ser	Ser	Met 215	Gly	Lys	Glu	Glu	Leu 220	Arg	His	Arg	Arg	Ala 225
Gln	Met	Pro	Thr	Pro 230	Lys	Ala	Ile	Asp	Cys 235	Arg	Lys	Cys	Phe	Gly 240

-----

PF-0459 US

Ala Pro Pro Glu Cys 245

# (2) INFORMATION FOR SEQ ID NO: 58:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 310 amino acids (B) TYPE: amino acid

  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: GBLANOT02
- (B) CLONE: 2530650

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 58:

Met	Leu	Leu	Pro	Gln 5	Leu	Cys	Trp	Leu	Pro 10	Leu	Leu	Ala	Gly	Leu 15
Leu	Pro	Pro	Val	Pro 20	Ala	Gln	Lys	Phe	Ser 25	Ala	Leu	Thr	Phe	Leu 30
_				35			Lys		40					45
				50			Cys		55					60
				65			Arg		70					75
Leu	Glu	Ile	Ala	80			Asn		85					90
		Glu	_	95	_		Gln		100					105
				110			Cys		115					Ser 120
Gln	Val	Gln	Cys	His 125			Thr		130					Thr 135
		Gly		140			Gly		145					Thr 150
				155			Asn		160					165
				170			Ala		175					180
				185			Asp		190					195
	_			200			Ser		205					210
				215			Gln		220					225
				230			Asp		235					240
				245			Pro		250					255
_	_			260			Val		265					270
_				275			Gln		280					285
Gln	Gly	Pro	Pro	Ser 290	Gln	Ser	Pro	Gly	Pro 295		Gln	Gly	Pro	Pro 300
Ala	Thr	Arg	Leu	Ser	Gly	Cys	Gln	Lys	Ala					

305

310

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 256 amino acids

(2) INFORMATION FOR SEQ ID NO: 59:

- (B) TYPE: amino acid
  (C) STRANDEDNESS: single
  (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: THYMNOT04
  - (B) CLONE: 2652271
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 59:

Met	Arg	Pro	Ala	Ala 5	Leu	Arg	Gly	Ala	Leu 10	Leu	Gly	Cys	Leu	Cys 15
Leu	Ala	Leu	Leu	Cys 20	Leu	Gly	Gly	Ala	Asp 25	Lys	Arg	Leu	Arg	Asp 30
				35				Met	40					45
		_		50				Asp	55					60
-	-			65				Pro	70					75
Asn	Arg	Ser	Trp	Pro 80				Glu	85					90
			Arg	95	_	_		Asp	100					105
				110				Glu	115					120
_				125				Asn	130					135
_	_			140				Glu	145					Val 150
		_		155				Ser	160					165
				170				Arg	175					180
Lys	Ile	Gln	Cys	Leu 185	Pro	Pro	Ser	Gln	Asp 190	Glu	Glu	Val	Gln	Thr 195
Ile	Gly	Gln	Ile	Glu 200	Leu	Cys	Leu	Thr	Lys 205	Gln	Asp	Gln	Gln	Leu 210
Gln	Asn	Cys	Thr	Glu 215	Pro	Gly	Glu	Gln	Pro 220	Ser	Pro	Lys	Gln	Glu 225
Val	Trp	Leu	Ala	Asn 230	Gly	Ala	Ala	Glu	Ser 235	Arg	Gly	Leu	Arg	Val 240
Cys	Glu	Asp	Gly	Pro 245	Val	Phe	Tyr	Pro	Pro 250	Pro	Lys	Lys	Thr	Lys 255
His														

- (2) INFORMATION FOR SEQ ID NO:
  - (i) SEQUENCE CHARACTERISTICS:

172

THE THE REAL PLANS STATE STATE

- (A) LENGTH: 160 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGTUT11
- (B) CLONE: 2746976
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 60:

```
Met Gln Phe Met Leu Leu Phe Ser Arg Gln Gly Lys Leu Arg Leu
                                      10
Gln Lys Trp Tyr Val Pro Leu Ser Asp Lys Glu Lys Arg Lys Ile
                 20
                                      25
Thr Arg Glu Leu Val Gln Thr Val Leu Ala Arg Lys Pro Lys Met
                                      40
                 35
Cys Ser Phe Leu Glu Trp Arg Asp Leu Lys Ile Val Tyr Lys Arg
                                      55
                 50
Tyr Ala Ser Leu Tyr Phe Cys Cys Ala Ile Glu Asp Gln Asp Asn
                                      70
                 65
Glu Leu Ile Thr Leu Glu Ile Ile His Arg Tyr Val Glu Leu Leu
                                                          90
                                      85
                 80
Asp Lys Tyr Phe Gly Ser Val Cys Glu Leu Asp Ile Ile Phe Asn
                                     100
                 95
Phe Glu Lys Ala Tyr Phe Ile Leu Asp Glu Phe Leu Leu Gly Gly
                                                         120
                                     115
                110
Glu Val Gln Glu Thr Ser Lys Lys Asn Val Leu Lys Ala Ile Glu
                                     130
                125
Gln Ala Asp Leu Leu Gln Glu Asp Ala Lys Glu Ala Glu Thr Pro
                                     145
                140
Arg Ser Val Leu Glu Glu Ile Gly Leu Thr
                155
```

#### (2) INFORMATION FOR SEQ ID NO: 61:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 341 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: THP1AZS08
- (B) CLONE: 2753496
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 61:

Met Lys Arg Ala Leu Gly Arg Arg Lys Gly Val Trp Leu Arg Leu 10 Arg Lys Ile Leu Phe Cys Val Leu Gly Leu Tyr Ile Ala Ile Pro 30 20 25 Phe Leu Ile Lys Leu Cys Pro Gly Ile Gln Ala Lys Leu Ile Phe 35 40 Leu Asn Phe Val Arg Val Pro Tyr Phe Ile Asp Leu Lys Lys Pro 60 55 50 Gln Asp Gln Gly Leu Asn His Thr Cys Asn Tyr Tyr Leu Gln Pro 70 Glu Glu Asp Val Thr Ile Gly Val Trp His Thr Val Pro Ala Val

				80					85					90
				95					100				Glu	105
Ala	Leu	Ala	Ser	Ser 110	His	Pro	Ile	Ile	Leu 115	Tyr	Leu	His	Gly	Asn 120
Ala	Gly	Thr	Arg	Gly 125	Gly	Asp	His	Arg	Val 130	Glu	Leu	Tyr	Lys	Val 135
Leu	Ser	Ser	Leu	Gly 140	Tyr	His	Val	Val	Thr 145	Phe	Asp	Tyr	Arg	Gly 150
Trp	Gly	Asp	Ser	Val 155	Gly	Thr	Pro	Ser	Glu 160	Arg	Gly	Met	Thr	Tyr 165
Asp	Ala	Leu	His	Val 170	Phe	Asp	Trp	Ile	Lys 175	Ala	Arg	Ser	Gly	Asp 180
Asn	Pro	Val	Tyr	Ile 185	Trp	Gly	His	Ser	Leu 190	Gly	Thr	Gly	Val	Ala 195
Thr	Asn	Leu	Val	Arg 200	Arg	Leu	Cys	Glu	Arg 205	Glu	Thr	Pro	Pro	Asp 210
Ala	Leu	Ile	Leu	Glu 215	Ser	Pro	Phe	Thr	Asn 220	Ile	Arg	Glu	Glu	Ala 225
				230					235				Gly	240
				245					250				Lys	255
				260					265				Leu	270
				275					280				Gly	285
				290					295				Asp	300
				305					310				Arg	315
				320					325		Ile	Leu	Arg	Glu 330
Phe	Leu	Gly	Lys	Ser 335	Glu	Pro	Glu	His	Gln 340	His				

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 430 amino acids (B) TYPE: amino acid

  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: OVARTUT03
- (B) CLONE: 2781553

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 62:

Met Ala Glu Gly Glu Asp Val Gly Trp Trp Arg Ser Trp Leu Gln 15 Gln Ser Tyr Gln Ala Val Lys Glu Lys Ser Ser Glu Ala Leu Glu 20 25 Phe Met Lys Arg Asp Leu Thr Glu Phe Thr Gln Val Val Gln His 35 40 Asp Thr Ala Cys Thr Ile Ala Ala Thr Ala Ser Val Val Lys Glu 50

62:

```
Lys Leu Ala Thr Glu Gly Ser Ser Gly Ala Thr Glu Lys Met Lys
                                     70
                 65
Lys Gly Leu Ser Asp Phe Leu Gly Val Ile Ser Asp Thr Phe Ala
                                     85
                 80
Pro Ser Pro Asp Lys Thr Ile Asp Cys Asp Val Ile Thr Leu Met
                                    100
                 95
Gly Thr Pro Ser Gly Thr Ala Glu Pro Tyr Asp Gly Thr Lys Ala
                                    115
                110
Arg Leu Tyr Ser Leu Gln Ser Asp Pro Ala Thr Tyr Cys Asn Glu
                                    130
                125
Pro Asp Gly Pro Pro Glu Leu Phe Asp Ala Trp Leu Ser Gln Phe
                                    145
                140
Cys Leu Glu Glu Lys Lys Gly Glu Ile Ser Glu Leu Leu Val Gly
                                    160
                155
Ser Pro Ser Ile Arg Ala Leu Tyr Thr Lys Met Val Pro Ala Ala
                                    175
                170
Val Ser His Ser Glu Phe Trp His Arg Tyr Phe Tyr Lys Val His
                                                         195
                185
                                    190
Gln Leu Glu Gln Glu Gln Ala Arg Arg Asp Ala Leu Lys Gln Arg
                200
                                    205
Ala Glu Gln Ser Ile Ser Glu Glu Pro Gly Trp Glu Glu Glu
                                    220
                215
Glu Glu Leu Met Gly Ile Ser Pro Ile Ser Pro Lys Glu Ala Lys
                                                         240
                230
                                    235
Val Pro Val Ala Lys Ile Ser Thr Phe Pro Glu Gly Glu Pro Gly
                                    250
                                                         255
                245
Pro Gln Ser Pro Cys Glu Glu Asn Leu Val Thr Ser Val Glu Pro
                                     265
                260
Pro Ala Glu Val Thr Pro Ser Glu Ser Ser Glu Ser Ile Ser Leu
                                    280
                275
Val Thr Gln Ile Ala Asn Pro Ala Thr Ala Pro Glu Ala Arg Val
                                                         300
                290
                                    295
Leu Pro Lys Asp Leu Ser Gln Lys Leu Leu Glu Ala Ser Leu Glu
                                     310
                                                         315
                305
Glu Gln Gly Leu Ala Val Asp Val Gly Glu Thr Gly Pro Ser Pro
                                                         330
                320
                                     325
Pro Ile His Ser Lys Pro Leu Thr Pro Ala Gly His Thr Gly Gly
                                     340
                                                         345
                335
Pro Glu Pro Arg Pro Pro Ala Arg Val Glu Thr Leu Arg Glu Glu
                350
                                     355
Ala Pro Thr Asp Leu Arg Val Phe Glu Leu Asn Ser Asp Ser Gly
                                                         375
                                     370
                365
Lys Ser Thr Pro Ser Asn Asn Gly Lys Lys Gly Ser Ser Thr Asp
                 380
                                     385
Ile Ser Glu Asp Trp Glu Lys Asp Phe Asp Leu Asp Met Thr Glu
                395
                                     400
Glu Glu Val Gln Met Ala Leu Ser Lys Val Asp Ala Ser Gly Glu
                                     415
                 410
Leu Glu Asp Val Glu Trp Glu Asp Trp Glu
```

### (2) INFORMATION FOR SEQ ID NO:

- 63:
- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 143 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

## (vii) IMMEDIATE SOURCE:

(A) LIBRARY: ADRETUTO6

(B) CLONE: 2821925

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 63:

Met Gly Pro Val Arg Leu Gly Ile Leu Leu Phe Leu Phe Leu Ala 10 Val His Glu Ala Trp Ala Gly Met Leu Lys Glu Glu Asp Asp Asp 25 30 20 Thr Glu Arg Leu Pro Ser Lys Cys Glu Val Cys Lys Leu Leu Ser 40 Thr Glu Leu Gln Ala Glu Leu Ser Arg Thr Gly Arg Ser Arg Glu 55 50 Val Leu Glu Leu Gly Gln Val Leu Asp Thr Gly Lys Arg Lys Arg 65 70 His Val Pro Tyr Ser Val Ser Glu Thr Arg Leu Glu Glu Ala Leu 85 80 Glu Asn Leu Cys Glu Arg Ile Leu Asp Tyr Ser Val His Ala Glu 100 95 Arg Lys Gly Ser Leu Arg Tyr Ala Lys Gly Gln Ser Gln Thr Met 115 110 Ala Thr Leu Lys Gly Leu Val Gln Lys Gly Val Lys Val Asp Leu 125 Gly Ile Pro Leu Glu Leu Leu Gly 140

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 301 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

## (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: UTRSTUT05
- (B) CLONE: 2879068

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 64:

Met Glu Asp Met Asn Glu Tyr Ser Asn Ile Glu Glu Phe Ala Glu 5 10 Gly Ser Lys Ile Asn Ala Ser Lys Asn Gln Gln Asp Asp Gly Lys 30 25 Met Phe Ile Gly Gly Leu Ser Trp Asp Thr Ser Lys Lys Asp Leu 40 35 Thr Glu Tyr Leu Ser Arg Phe Gly Glu Val Val Asp Cys Thr Ile 50 55 Lys Thr Asp Pro Val Thr Gly Arg Ser Arg Gly Phe Gly Phe Val 70 65 Leu Phe Lys Asp Ala Ala Ser Val Asp Lys Val Leu Glu Leu Lys 85 Glu His Lys Leu Asp Gly Lys Leu Ile Asp Pro Lys Arg Ala Lys 105 100 95 Ala Leu Lys Gly Lys Glu Pro Pro Lys Lys Val Phe Val Gly Gly 110 115 Leu Ser Pro Asp Thr Ser Glu Glu Gln Ile Lys Glu Tyr Phe Gly

				125					130					135
Ala	Phe	Gly	Glu	Ile 140	Glu	Asn	Ile	Glu	Leu 145	Pro	Met	Asp	Thr	Lys 150
Thr	Asn	Glu	Arg	Arg 155	Gly	Phe	Cys	Phe	Ile 160	Thr	Tyr	Thr	Asp	Glu 165
Glu	Pro	Val	Lys	Lys 170	Leu	Leu	Glu	Ser	Arg 175	Tyr	His	Gln	Ile	Gly 180
Ser	Gly	Lys	Cys	Glu 185	Ile	Lys	Val	Ala	Gln 190	Pro	Lys	Glu	Val	Tyr 195
Arg	Gln	Gln	Gln	Gln 200	Gln	Gln	Lys	Gly	Gly 205	Arg	Gly	Ala	Ala	Ala 210
Gly	Gly	Arg	Gly	Gly 215	Thr	Arg	Gly	Arg	Gly 220	Arg	Gly	Gln	Gly	Gln 225
Asn	Trp	Asn	Gln	Gly 230	Phe	Asn	Asn	Tyr	Tyr 235	Asp	Gln	Gly	Tyr	Gly 240
Asn	Tyr	Asn	Ser	Ala 245	Tyr	Gly	Gly	Asp	Gln 250	Asn	Tyr	Ser	Gly	Tyr 255
Gly	Gly	Tyr	Asp	Tyr 260	Thr	Gly	Tyr	Asn	Tyr 265	Gly	Asn	Tyr	Gly	Tyr 270
Gly	Gln	Gly	Tyr	Ala 275	Asp	Tyr	Ser	Gly	Gln 280	Gln	Ser	Thr	Tyr	Gly 285
Lys	Ala	Ser	Arg	Gly 290	Gly	Gly	Asn	His	Gln 295	Asn	Asn	Tyr	Gln	Pro 300
Tyr														

## (2) INFORMATION FOR SEQ ID NO: 65:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 233 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: SINJNOT02 (B) CLONE: 2886757

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 65:

Met	Gly	Glu	Pro	Gln 5	Gln	Val	Ser	Ala	Leu 10	Pro	Pro	Pro	Pro	Met 15
Gln	Tyr	Ile	Lys	Glu 20	Tyr	Thr	Asp	Glu	Asn 25	Ile	Gln	Glu	Gly	Leu 30
Ala	Pro	Lys	Pro	Pro 35	Pro	Pro	Ile	Lys	Asp 40	Ser	Tyr	Met	Met	Phe 45
Gly	Asn	Gln	Phe	Gln 50	Cys	Asp	Asp	Leu	Ile 55	Ile	Arg	Pro	Leu	Glu 60
Ser	Gln	Gly	Ile	Glu 65	Arg	Leu	His	Pro	Met 70	Gln	Phe	Asp	His	Lys 75
Lys	Glu	Leu	Arg	Lys 80	Leu	Asn	Met	Ser	Ile 85	Leu	Ile	Asn	Phe	Leu 90
Asp	Leu	Leu	Asp	Ile 95	Leu	Ile	Arg	Ser	Pro 100	Gly	Ser	Ile	Lys	Arg 105
Glu	Glu	Lys	Leu	Glu 110	Asp	Leu	Lys	Leu	Leu 115	Phe	Val	His	Val	His 120
His	Leu	Ile	Asn	Glu 125	Tyr	Arg	Pro	His	Gln 130	Ala	Arg	Glu	Thr	Leu 135
Arg	Val	Met	Met	Glu 140	Val	Gln	Lys	Arg	Gln 145	Arg	Leu	Glu	Thr	Ala 150

```
Glu Arg Phe Gln Lys His Leu Glu Arg Val Ile Glu Met Ile Gln
                155
                                   160
Asn Cys Leu Ala Ser Leu Pro Asp Asp Leu Pro His Ser Glu Ala
                                    175
                170
Gly Met Arg Val Lys Thr Glu Pro Met Asp Ala Asp Asp Ser Asn
                185
                                    190
Asn Cys Thr Gly Gln Asn Glu His Gln Arg Glu Asn Ser Gly His
                                   205
                200
Arg Arg Asp Gln Ile Ile Glu Lys Asp Ala Ala Leu Cys Val Leu
                                                        225
                215
Ile Asp Glu Met Asn Glu Arg Pro
                230
```

66:

#### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 354 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: SCORNOT04
- (B) CLONE: 2964329

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 66:

Met	Ala	Gly	Ala	Gly 5	Ala	Gly	Ala	Gly	Ala 10	Arg	Gly	Gly	Ala	Ala 15
Ala	Gly	Val	Glu	Ala 20	Arg	Ala	Arg	Asp	Pro 25	Pro	Pro	Ala	His	Arg 30
Ala	His	Pro	Arg	His 35	Pro	Arg	Pro	Ala	Ala 40	Gln	Pro	Ser	Ala	Arg 45
Arg	Met	Asp	Gly	Gly 50	Ser	Gly	Gly	Leu	Gly 55	Ser	Gly	Asp	Asn	Ala 60
Pro	Thr	Thr	Glu	Ala 65	Leu	Phe	Val	Ala	Leu 70	Gly	Ala	Gly	Val	Thr 75
Ala	Leu	Ser	His	Pro 80	Leu	Leu	Tyr	Val	Lys 85	Leu	Leu	Ile	Gln	Val 90
Gly	His	Glu	Pro	Met 95	Pro	Pro	Thr	Leu	Gly 100	Thr	Asn	Val	Leu	Gly 105
Arg	Lys	Val	Leu	Tyr 110	Leu	Pro	Ser	Phe	Phe 115	Thr	Tyr	Ala	Lys	Tyr 120
Ile	Val	Gln	Val	Asp 125	Gly	Lys	Ile	Gly	Leu 130	Phe	Arg	Gly	Leu	Ser 135
Pro	Arg	Leu	Met	Ser 140	Asn	Ala	Leu	Ser	Thr 145	Val	Thr	Arg	Gly	Ser 150
Met	Lys	Lys	Val	Phe 155	Pro	Pro	Asp	Glu	Ile 160	Glu	Gln	Val	Ser	Asn 165
Lys	Asp	Asp	Met		Thr	Ser	Leu	Lys	Lys 175	Val	Val	Lys	Glu	Thr 180
Ser	Tyr	Glu	Met		Met	Gln	Cys	Val	Ser 190	Arg	Met	Leu	Ala	His 195
Pro	Leu	His	Val		Ser	Met	Arg	Cys	Met 205	Val	Gln	Phe	Val	Gly 210
Arg	Glu	Ala	Lys		Ser	Gly	Val	Leu		Ser	Ile	Gly	Lys	Ile 225
Phe	Lys	Glu	Glu		Leu	Leu	Gly	Phe		Val	Gly	Leu	Ile	Pro

				230					235					240
His	Leu	Leu	Gly	Asp 245	Val	Val	Phe	Leu	Trp 250	Gly	Cys	Asn	Leu	Leu 255
Ala	His	Phe	Ile	Asn 260	Ala	Tyr	Leu	Val	Asp 265	Asp	Ser	Phe	Ser	Gln 270
Ala	Leu	Ala	Ile	Arg 275	Ser	Tyr	Thr	Lys	Phe 280	Val	Met	Gly	Ile	Ala 285
Val	Ser	Met	Leu	Thr 290	Tyr	Pro	Phe	Leu	Leu 295	Val	Gly	Asp	Leu	Met 300
Ala	Val	Asn	Asn	Cys 305	Gly	Leu	Gln	Ala	Gly 310	Leu	Pro	Pro	Tyr	Ser 315
Pro	Val	Phe	Lys	Ser 320	Trp	Ile	His	Cys	Trp 325	Lys	Tyr	Leu	Ser	Val 330
Gln	Gly	Gln	Leu	Phe 335	Arg	Gly	Ser	Ser	Leu 340	Leu	Phe	Arg	Arg	Val 345
Ser	Ser	Gly	Ser	Cys 350	Phe	Ala	Leu	Glu						

## (2) INFORMATION FOR SEQ ID NO: 67:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 235 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
    (D) TOPOLOGY: linear

## (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: SCORNOT04
- (B) CLONE: 2965248

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 67:

Met	Ala	Ser	Thr	Ile 5	Ser	Ala	Tyr	Lys	Glu 10	Lys	Met	Lys	Glu	Leu 15
Ser	Val	Leu	Ser	_	Ile	Cys	Ser	Cys	Phe 25	Tyr	Thr	Gln	Pro	His 30
Pro	Asn	Thr	Val	Tyr 35	Gln	Tyr	Gly	Asp	Met 40	Glu	Val	Lys	Gln	Leu 45
Asp	Lys	Arg	Ala	Ser 50	Gly	Gln	Ser	Phe	Glu 55	Val	Ile	Leu	Lys	Ser 60
Pro	Ser	Asp	Leu	Ser 65	Pro	Glu	Ser	Pro	Met 70	Leu	Ser	Ser	Pro	Pro 75
Lys	Lys	Lys	Asp	Thr 80	Ser	Leu	Glu	Glu	Leu 85	Gln	Lys	Arg	Leu	Glu 90
Ala	Ala	Glu	Glu	Arg 95	Arg	Lys	Thr	Gln	Glu 100	Ala	Gln	Val	Leu	Lys 105
Gln	Leu	Ala	Asp	Gly 110	Ala	Ser	Thr	Ser	Ala 115	Arg	Cys	Cys	Thr	Arg 120
Arg	Trp	Arg	Arg	Ile 125	Thr	Thr	Ser	Ala	Ala 130	Arg	Arg	Arg	Arg	Ser 135
Ser	Thr	Thr	Arg	Trp 140	Ser	Ser	Ala	Arg	Arg 145	Ser	Ala	Arg	His	Thr 150
Trp	Pro	His	Cys	Ala 155	Ser	Gly	Cys	Ala	Arg 160	Arg	Ser	Cys	Thr	Arg 165
Pro	Arg	Cys	Ala	Gly 170	Thr	Arg	Ser	Ser	Glu 175	Lys	Arg	Cys	Arg	Ala 180
Lys	Gly	Pro	Gly	Arg 185	Ala	Ala	Pro	Ile	Leu 190	Arg	Arg	Asn	Thr	Phe 195

```
Gly Phe Trp Phe Cys Phe Val His Leu Cys Leu Asp Ala Thr Phe 200 Val Pro Pro Gln Pro Pro Ala Ser Cys Phe Ser Ser Ala Leu Ser Arg Pro Ala Leu Ser Ser Trp 230 Val Pro Ser Ser 235 Val Pro Pro Ala Leu Ser Ser Trp 235 Variable.
```

### (2) INFORMATION FOR SEQ ID NO: 68:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 221 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: TLYMNOT06
- (B) CLONE: 3000534
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 68:

			Ala	5					10					15
			Leu	20					25					30
	_		Glu	35					40					45
			His	50					55					60
			Ser	65					70					75
_	_		Asn	80					85					90
_			Arg	95					100					105
			Val	110					115					120
			Leu	125					130					135
			Gly	140					145					150
_			Trp	155					160					165
			Val	170					175					180
			Gly	185					190					195
Thr	His	Asn	Thr	Trp 200	Lys	Ala	Met	Glu	Gly 205	Ile	Phe	Ile	Lys	Pro 210
Ser	Val	Glu	Pro	Ser 215	Ala	Gly	His	Asp	Glu 220	Leu				

- (2) INFORMATION FOR SEQ ID NO: 69:
  - (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 483 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: HEAANOT01
- (B) CLONE: 3046870

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 69:

```
Met Lys Ala Phe His Thr Phe Cys Val Val Leu Leu Val Phe Gly
                                      10
Ser Val Ser Glu Ala Lys Phe Asp Asp Phe Glu Asp Glu Glu Asp
                                      25
                 20
Ile Val Glu Tyr Asp Asp Asn Asp Phe Ala Glu Phe Glu Asp Val
                                      40
                 35
Met Glu Asp Ser Val Thr Glu Ser Pro Gln Arg Val Ile Ile Thr
                                      55
                 50
Glu Asp Asp Glu Asp Glu Thr Thr Val Glu Leu Glu Gly Gln Asp
                                                          75
                                      70
                  65
Glu Asn Gln Glu Gly Asp Phe Glu Asp Ala Asp Thr Gln Glu Gly
                                                          90
                                      85
                 80
Asp Thr Glu Ser Glu Pro Tyr Asp Asp Glu Glu Phe Glu Gly Tyr
                                                          105
                                     100
                 95
Glu Asp Lys Pro Asp Thr Ser Ser Ser Lys Asn Lys Asp Pro Ile
                110
                                     115
Thr Ile Val Asp Val Pro Ala His Leu Gln Asn Ser Trp Glu Ser
                                     130
                 125
Tyr Tyr Leu Glu Ile Leu Met Val Thr Gly Leu Leu Ala Tyr Ile
                                                          150
                                     145
                140
Met Asn Tyr Ile Ile Gly Lys Asn Lys Asn Ser Arg Leu Ala Gln
                                     160
                 155
Ala Trp Phe Asn Thr His Arg Glu Leu Leu Glu Ser Asn Phe Thr
                                     175
                                                          180
                 170
Leu Val Gly Asp Asp Gly Thr Asn Lys Glu Ala Thr Ser Thr Gly
                                     190
                                                          195
                 185
Lys Leu Asn Gln Glu Asn Glu His Ile Tyr Asn Leu Trp Cys Ser
                                                          210
                                     205
                 200
Gly Arg Val Cys Cys Glu Gly Met Leu Ile Gln Leu Arg Phe Leu
                                     220
                 215
Lys Arg Gln Asp Leu Leu Asn Val Leu Ala Arg Met Met Arg Pro
                 230
                                      235
Val Ser Asp Gln Val Gln Ile Lys Val Thr Met Asn Asp Glu Asp
                                      250
                 245
Met Asp Thr Tyr Val Phe Ala Val Gly Thr Arg Lys Ala Leu Val
                                                          270
                                      265
                 260
Arg Leu Gln Lys Glu Met Gln Asp Leu Ser Glu Phe Cys Ser Asp
                                     280
                                                          285
                 275
 Lys Pro Lys Ser Gly Ala Lys Tyr Gly Leu Pro Asp Ser Leu Ala
                                                          300
                                      295
                 290
 Ile Leu Ser Glu Met Gly Glu Val Thr Asp Gly Met Met Asp Thr
                                      310
                 305
 Lys Met Val His Phe Leu Thr His Tyr Ala Asp Lys Ile Glu Ser
                                                          330
                                      325
                 320
 Val His Phe Ser Asp Gln Phe Ser Gly Pro Lys Ile Met Gln Glu
                                      340
                 335
 Glu Gly Gln Pro Leu Lys Leu Pro Asp Thr Lys Arg Thr Leu Leu
                                      355
                 350
 Phe Thr Phe Asn Val Pro Gly Ser Gly Asn Thr Tyr Pro Lys Asp
                                      370
                 365
```

```
Met Glu Ala Leu Leu Pro Leu Met Asn Met Val Ile Tyr Ser Ile
                380
                                     385
Asp Lys Ala Lys Lys Phe Arg Leu Asn Arg Glu Gly Lys Gln Lys
                                     400
                395
Ala Asp Lys Asn Arg Ala Arg Val Glu Glu Asn Phe Leu Lys Leu
                410
                                     415
Thr His Val Gln Arq Gln Glu Ala Ala Gln Ser Arg Arg Glu Glu
                                                          435
                                     430
                425
Lys Lys Arg Ala Glu Lys Glu Arg Ile Met Asn Glu Glu Asp Pro
                440
                                                          450
Glu Lys Gln Arg Arg Leu Glu Glu Ala Ala Leu Arg Arg Glu Gln
                                     460
                455
Lys Lys Leu Glu Lys Lys Gln Met Lys Met Lys Gln Ile Lys Val
                                                          480
                470
Lys Ala Met
```

### (2) INFORMATION FOR SEQ ID NO:

70:

# (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 371 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: PONSAZT01
- (B) CLONE: 3057669

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 70:

Met	Asp	His	Glu	Asp 5	Ile	Ser	Glu	Ser	Val 10	Asp	Ala	Ala	Tyr	Asn 15
Leu	Gln	Asp	Ser	Cys 20	Leu	Thr	Asp	Cys	Asp 25	Val	Glu	Asp	Gly	Thr 30
Met	Asp	Gly	Asn	Asp 35	Glu	Gly	His	Ser	Phe 40	Glu	Leu	Cys	Pro	Ser 45
Glu	Ala	Ser	Pro	Tyr 50	Val	Arg	Ser	Arg	Glu 55	Arg	Thr	Ser	Ser	Ser 60
Ile	Val	Phe	Glu	Asp 65	Ser	Gly	Cys	Asp	Asn 70	Ala	Ser	Ser	Lys	Glu 75
Glu	Pro	Lys	Thr	Asn 80	Arg	Leu	His	Ile	Gly 85	Asn	His	Суѕ	Ala	Asn 90
Lys	Leu	Thr	Ala	Phe 95	Lys	Pro	Thr	Ser	Ser 100	Lys	Ser	Ser	Ser	Glu 105
Ala	Thr	Leu	Ser	Ile 110	Ser	Pro	Pro	Arg	Pro 115	Thr	Thr	Leu	Ser	Leu 120
Asp	Leu	Thr	Lys	Asn 125	Thr	Thr	Glu	Lys	Leu 130	Gln	Pro	Ser	Ser	Pro 135
Lys	Val	Tyr	Leu	Tyr 140	Ile	Gln	Met	Gln	Leu 145	Суѕ	Arg	Lys	Glu	Asn 150
Leu	Lys	Asp	Trp	Met 155	Asn	Gly	Arg	Cys	Thr 160	Ile	Glu	Glu	Arg	Glu 165
Arg	Ser	Val	Cys	Leu 170	His	Ile	Phe	Leu	Gln 175	Ile	Ala	Glu	Ala	Val 180
Glu	Phe	Leu	His	Ser 185	Lys	Gly	Leu	Met	His 190	Arg	Asp	Leu	Lys	Pro 195
Ser	Asn	Ile	Phe	Phe 200	Thr	Met	Asp	Asp	Val 205	Val	Lys	Val	Gly	Asp 210
Phe	Gly	Leu	Val	Thr	Ala	Met	Asp	Gln	Asp	Glu	Glu	Glu	Gln	Thr

				215					220					225
Val	Leu	Thr	Pro	Met	Pro	Ala	Tyr	Ala	Arg	His	Thr	Gly	Gln	Val
				230					235				_	240
Gly	Thr	Lys	Leu	Tyr	Met	Ser	Pro	Glu	Gln	Ile	His	GTA	Asn	Ser
				245					250		_			255
Tyr	Ser	His	Lys	Val	Asp	Ile	Phe	Ser		Gly	Leu	TTe	Leu	Phe
				260					265		_		_	270
Glu	Leu	Leu	Tyr	Pro	Phe	Ser	Thr	Gln	Met	Glu	Arg	Val	Arg	Thr
				275					280			_,	<b>-</b>	285
Leu	Thr	Asp	Val	Arg	Asn	Leu	Lys	Phe	Pro	Pro	Leu	Phe	Thr	Gln
				290					295			_		300
Lys	Tyr	Pro	Cys	Glu	Tyr	Val	Met	Val		Asp	Met	Leu	Ser	Pro
				305				_	310			<b>~</b> 1	-	315
Ser	Pro	Met	Glu		Pro	Glu	Ala	Ile	Asn	Ile	ııe	Glu	Asn	Ala
				320				_	325		3		-	330
Val	Phe	Glu	Asp	Leu	Asp	Phe	Pro	Gly	Lys	Thr	Val	Leu	Arg	Gln
				335					340	_	•	~	_	345
Arg	Ser	Arg	Ser		Ser	Ser	Ser	Gly	Thr	Lys	His	Ser	Arg	GIN
				350					355					360
Ser	Asn	Asn	Ser	His	Ser	Pro	Leu	Pro		Asn				
				365					370					

# (2) INFORMATION FOR SEQ ID NO: 71:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 402 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

## (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: HEAONOT03 (B) CLONE: 3088178

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 71:

Met	Met	Asn	Asn	Arg 5	Phe	Arg	Lys	Asp	Met 10	Met	Lys	Asn	Ala	Ser 15
Glu	Ser	Lys	Leu	Ser 20	Lys	Asp	Asn	Leu	Lys 25	Lys	Arg	Leu	Lys	Glu 30
Glu	Phe	Gln	His	Ala 35	Met	Gly	Gly	Val	Pro 40	Ala	Trp	Ala	Glu	Thr 45
	_			50					55				Asp	Glu 60
_				65					70	Ile				Thr 75
Ser	Leu	Pro	Arg	Gly 80	Ile	Leu	Lys	Met	Lys 85	Asn	Cys	Gln	His	Ala 90
Asn	Ala	Glu	Arg	Pro 95	Thr	Val	Ala	Arg	Ile 100	Ser	Ser	Val	Gln	Phe 105
His	Pro	Gly	Ala	Gln 110	Ile	Val	Met	Val	Ala 115	Gly	Leu	Asp	Asn	Ala 120
Val	Ser	Leu	Phe	Gln 125	Val	Asp	Gly	Lys	Thr 130	Asn	Pro	Lys	Ile	Gln 135
Ser	Ile	Tyr	Leu	Glu 140	Arg	Phe	Pro	Ile	Phe 145	Lys	Ala	Cys	Phe	Ser 150
Ala	Asn	Gly	Glu	Glu 155	Val	Leu	Ala	Thr	Ser 160	Thr	His	Ser	Lys	Val 165

```
Leu Tyr Val Tyr Asp Met Leu Ala Gly Lys Leu Ile Pro Val His
                                     175
                170
Gln Val Arg Gly Leu Lys Glu Lys Ile Val Arg Ser Phe Glu Val
                                     190
                185
Ser Pro Asp Gly Ser Phe Leu Leu Ile Asn Gly Ile Ala Gly Tyr
                                                         210
                200
                                     205
Leu His Leu Leu Ala Met Lys Thr Lys Glu Leu Ile Gly Ser Met
                                     220
                215
Lys Ile Asn Gly Arg Val Ala Ala Ser Thr Phe Ser Ser Asp Ser
                230
                                     235
Lys Lys Val Tyr Ala Ser Ser Gly Asp Gly Glu Val Tyr Val Trp
                                     250
                245
Asp Val Asn Ser Arg Lys Cys Leu Asn Arg Phe Val Asp Glu Gly
                                     265
                                                         270
                260
Ser Leu Tyr Gly Leu Ser Ile Ala Thr Ser Arg Asn Gly Gln Tyr
                                     280
                                                         285
                275
Val Ala Cys Gly Ser Asn Cys Gly Val Val Asn Ile Tyr Asn Gln
                                                         300
                                     295
                290
Asp Ser Cys Leu Gln Glu Thr Asn Pro Lys Pro Ile Lys Ala Ile
                                     310
                                                         315
                305
Met Asn Leu Val Thr Gly Val Thr Ser Leu Thr Phe Asn Pro Thr
                                     325
                                                         330
                320
Thr Glu Ile Leu Ala Ile Ala Ser Glu Lys Met Lys Glu Ala Val
                                                         345
                335
                                     340
Arg Leu Val His Leu Pro Ser Cys Thr Val Phe Ser Asn Phe Pro
                                     355
                                                         360
                350
Val Ile Lys Asn Lys Asn Ile Ser His Val His Thr Met Asp Phe
                                     370
                 365
Ser Pro Arg Ser Gly Tyr Phe Ala Leu Gly Asn Glu Lys Gly Lys
                                     385
                                                          390
                 380
Ala Leu Met Tyr Arg Leu His His Tyr Ser Asp Phe
                 395
```

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 640 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

## (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRSTNOT19
- (B) CLONE: 3094321

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 72:

Met Ala Leu Ser Arg Gly Leu Pro Arg Glu Leu Ala Glu Ala Val 10 Ala Gly Gly Arg Val Leu Val Val Gly Ala Gly Gly Ile Gly Cys 25 30 20 Glu Leu Leu Lys Asn Leu Val Leu Thr Gly Phe Ser His Ile Asp 45 35 40 Leu Ile Asp Leu Asp Thr Ile Asp Val Ser Asn Leu Asn Arg Gln 55 50 Phe Leu Phe Gln Lys Lys His Val Gly Arg Ser Lys Ala Gln Val 70 Ala Lys Glu Ser Val Leu Gln Phe Tyr Pro Lys Ala Asn Ile Val

				80					85					90
Ala	Tyr	His	Asp		Ile	Met	Asn	Pro		Tyr	Asn	Val	Glu	
Phe	Arg	Gln	Phe	Ile 110	Leu	Val	Met	Asn	Ala 115	Leu	Asp	Asn	Arg	Ala 120
Ala	Arg	Asn	His		Asn	Arg	Met	Cys		Ala	Ala	Asp	Val	Pro 135
Leu	Ile	Glu	Ser	Gly 140	Thr	Ala	Gly	Tyr	Leu 145	Gly	Gln	Val	Thr	Thr 150
Ile	Lys	Lys	Gly		Thr	Glu	Cys	Tyr		Cys	His	Pro	Lys	
Thr	Gln	Arg	Thr		Pro	Gly	Cys	Thr	Ile 175	Arg	Asn	Thr	Pro	Ser 180
Glu	Pro	Ile	His	Cys 185	Ile	Val	Trp	Ala		Tyr	Leu	Phe	Asn	
Leu	Phe	Gly	Glu		Asp	Ala	Asp	Gln		Val	Ser	Pro	Asp	
Ala	Asp	Pro	Glu		Ala	Trp	Glu	Pro		Glu	Ala	Glu	Ala	Arg 225
Ala	Arg	Ala	Ser		Glu	Asp	Gly	Asp		Lys	Arg	Ile	Ser	Thr 240
Lys	Glu	Trp	Ala		Ser	Thr	Gly	Tyr	Asp 250	Pro	Val	Lys	Leu	Phe 255
Thr	Lys	Leu	Phe		Asp	Asp	Ile	Arg	Tyr 265	Leu	Leu	Thr	Met	Asp 270
Lys	Leu	Trp	Arg	Lys 275	Arg	Lys	Pro	Pro	Val 280	Pro	Leu	Asp	Trp	Ala 285
Glu	Val	Gln	Ser	Gln 290	Gly	Glu	Glu	Thr	Asn 295	Ala	Ser	Asp	Gln	Gln 300
			Gln	305					310					315
Lys	Ser	Tyr	Ala	Arg 320	Leu	Phe	Ser	Lys	Ser 325	Ile	Glu	Thr	Leu	Arg 330
Val	His	Leu	Ala	Glu 335	Lys	Gly	Asp	Gly	Ala 340	Glu	Leu	Ile	Trp	Asp 345
_	_		Pro	350					355					360
			His	365					370			,		375
Ile	Lys	Ser	Met	Ala 380	Gly	Asn	Ile	Ile	Pro 385	Ala	Ile	Ala	Thr	Thr 390
			Ile	395					400					405
Leu	Ser	Gly	Lys	Ile 410		Gln	. Cys	Arg	Thr 415	Ile	Phe	Leu	Asn	Lys 420
				425					Val 430	Pro				Asp 435
Pro	Pro	Asn	Pro	Asn 440		Tyr	· Val	Cys	Ala 445		Lys	Pro	Glu	Val 450
Thr	Val	Arg	Leu	Asn 455		His	Lys	Val	Thr 460		Leu	Thr	Leu	Gln 465
Asp	Lys	Ile	Val		Glu	Lys	Phe	Ala	Met 475		Ala	Pro	Asp	Val 480
Gln	Ile	Glu	Asp	Gly 485		Gly	Thr	Ile	Leu 490	Ile	Ser	Ser	Glu	Glu 495
Gly	Glu	Thr	Glu		Asn	Asn	His	Lys	Lys 505		Ser	Glu	Phe	Gly 510
Ile	Arg	Asn	Gly	Ser 515		Leu	ı Gln	Ala	Asp 520		Phe	Leu	Gln	Asp 525
Tyr	Thr	Leu	Leu		Asn	ı Ile	e Leu	His	Ser 535	Glu	Asp	Leu	Gly	Lys 540

```
Asp Val Glu Phe Glu Val Val Gly Asp Ala Pro Glu Lys Val Gly
                                 550 555
               545
Pro Lys Gln Ala Glu Asp Ala Ala Lys Ser Ile Thr Asn Gly Ser
               560
                                  565
Asp Asp Gly Ala Gln Pro Ser Thr Ser Thr Ala Gln Glu Gln Asp
               575
                                  580
Asp Val Leu Ile Val Asp Ser Asp Glu Glu Asp Ser Ser Asn Asn
                                   595
               590
Ala Asp Val Ser Glu Glu Glu Arg Ser Arg Lys Arg Lys Leu Asp
               605
                                   610
Glu Lys Glu Asn Leu Ser Ala Lys Arg Ser Arg Ile Glu Gln Lys
                                   625
               620
Glu Glu Leu Asp Asp Val Ile Ala Leu Asp
               635
```

### (2) INFORMATION FOR SEQ ID NO:

73:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 237 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGTUT13
- (B) CLONE: 3115936

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 73:

Met	Asp	Lys	Ile	Leu 5	Asn	Val	Glu	Glu	Thr 10	Tyr	Leu	Thr	Val	Leu 15
Val	Lys	Ile	Gly	Pro 20	Gly	Phe	His	Thr	Arg 25	Glu	Cys	Phe	Leu	Leu 30
Lys	Ser	Ile	Leu	Cys 35	Phe	Ser	Pro	Ser	Tyr 40	Arg	Met	Ser	Glu	Gly 45
Asp	Ser	Val	Gly	Glu 50	Ser	Val	His	Gly	Lys 55	Pro	Ser	Val	Val	Tyr 60
Arg	Phe	Phe	Thr	Arg 65	Leu	Gly	Gln	Ile	Tyr 70	Gln	Ser	Trp	Leu	Asp 75
Lys	Ser	Thr	Pro	Tyr 80	Thr	Ala	Val	Arg	Trp 85	Val	Val	Thr	Leu	Gly 90
			Val	95					100					105
Tyr	Ile	Val	Thr	Tyr 110	Ala	Leu	Gly	Ile	Tyr 115	His	Leu	Asn	Leu	Phe 120
Ile	Ala	Phe	Leu	Ser 125	Pro	Lys	Val	Asp	Pro 130	Ser	Leu	Met	Glu	Asp 135
Ser	Asp	Asp	Gly	Pro 140	Ser	Leu	Pro	Thr	Lys 145	Gln	Asn	Glu	Glu	Phe 150
Arg	Pro	Phe	Ile	Arg 155	Arg	Leu	Pro	Glu	Phe 160	Lys	Phe	Trp	His	Ala 165
Ala	Thr	Lys	Gly	Ile 170	Leu	Val	Ala	Met	Val 175	Cys	Thr	Phe	Phe	Asp 180
Ala	Phe	Asn	Val	Pro 185	Val	Phe	Trp	Pro	Ile 190	Leu	Val	Met	Tyr	Phe 195
Ile	Met	Leu	Phe	Cys 200	Ile	Thr	Met	Lys	Arg 205	Gln	Ile	Lys	His	Met 210
Ile	Lys	Tyr	Arg	Tyr	Ile	Pro	Phe	Thr	His	Gly	Lys	Arg	Arg	Tyr

215 220 225
Arg Gly Lys Glu Asp Ala Gly Lys Ala Phe Ala Ser
230 235

### (2) INFORMATION FOR SEQ ID NO: 74

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 432 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

## (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGTUT13
- (B) CLONE: 3116522

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 74:

```
Met Asp Ala Arg Trp Trp Ala Val Val Leu Ala Ala Phe Pro
                                     10
Ser Leu Gly Ala Gly Gly Glu Thr Pro Glu Ala Pro Pro Glu Ser
                                     25
                 20
Trp Thr Gln Leu Trp Phe Phe Arg Phe Val Val Asn Ala Ala Gly
                                     40
                 35
Tyr Ala Ser Phe Met Val Pro Gly Tyr Leu Leu Val Gln Tyr Phe
                                     55
                 50
Arg Arg Lys Asn Tyr Leu Glu Thr Gly Arg Gly Leu Cys Phe Pro
                                     70
                 65
Leu Val Lys Ala Cys Val Phe Gly Asn Glu Pro Lys Ala Ser Asp
                 80
                                     85
Glu Val Pro Leu Ala Pro Arg Thr Glu Ala Ala Glu Thr Thr Pro
                 95
                                    100
Met Trp Gln Ala Leu Lys Leu Leu Phe Cys Ala Thr Gly Leu Gln
                                     115
                110
Val Ser Tyr Leu Thr Trp Gly Val Leu Gln Glu Arg Val Met Thr
                                    130
                125
Arg Ser Tyr Gly Ala Thr Ala Thr Ser Pro Gly Glu Arg Phe Thr
                140
                                    145
Asp Ser Gln Phe Leu Val Leu Met Asn Arg Val Leu Ala Leu Ile
                                     160
                155
Val Ala Gly Leu Ser Cys Val Leu Cys Lys Gln Pro Arg His Gly
                                     175
                170
Ala Pro Met Tyr Arg Tyr Ser Phe Ala Ser Leu Ser Asn Val Leu
                                     190
                185
Ser Ser Trp Cys Gln Tyr Glu Ala Leu Lys Phe Val Ser Phe Pro
                 200
                                     205
Thr Gln Val Leu Ala Lys Ala Ser Lys Val Ile Pro Val Met Leu
                                     220
                                                         225
                215
Met Gly Lys Leu Val Ser Arg Arg Ser Tyr Glu His Trp Glu Tyr
                 230
                                     235
Leu Thr Ala Thr Leu Ile Ser Ile Gly Val Ser Met Phe Leu Leu
                                     250
                 245
Ser Ser Gly Pro Glu Pro Arg Ser Ser Pro Ala Thr Thr Leu Ser
                260
                                     265
Gly Leu Ile Leu Leu Ala Gly Tyr Ile Ala Phe Asp Ser Phe Thr
                                     280
                275
Ser Asn Trp Gln Asp Ala Leu Phe Ala Tyr Lys Met Ser Ser Val
                                     295
```

```
Gln Met Met Phe Gly Val Asn Phe Phe Ser Cys Leu Phe Thr Val
               305
                                   310
Gly Ser Leu Leu Glu Gln Gly Ala Leu Leu Glu Gly Thr Arg Phe
                                    325
               320
Met Gly Arg His Ser Glu Phe Ala Ala His Ala Leu Leu Ser
                335
                                   340
Ile Cys Ser Ala Cys Gly Gln Leu Phe Ile Phe Tyr Thr Ile Gly
                                    355
                350
Gln Phe Gly Ala Ala Val Phe Thr Ile Ile Met Thr Leu Arg Gln
                365
                                    370
Ala Phe Ala Ile Leu Leu Ser Cys Leu Leu Tyr Gly His Thr Val
                                    385
                380
Thr Val Val Gly Gly Leu Gly Val Ala Val Val Phe Ala Ala Leu
                                    400
                395
Leu Leu Arg Val Tyr Ala Arg Gly Arg Leu Lys Gln Arg Gly Lys
                                   415
               410
Lys Ala Val Pro Val Glu Ser Pro Val Gln Lys Val
                425
```

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 252 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
    (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGTUT13
- (B) CLONE: 3117184

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 75:

N/ - +	Com	Dho	Pro	Dro	Uic	Tou	Aen	Δrα	Pro	Pro	Met	Glv	Tle	Pro
				5					10					15
			Pro	20					25					30
Pro	Pro	Val	Pro	Pro 35	Gly	Thr	Pro	Met	Ile 40	Pro	Val	Pro	Met	Ser 45
Ile	Met	Ala	Pro	Ala 50	Pro	Thr	Val	Leu	Val 55	Pro	Thr	Val	Ser	Met 60
Val	Gly	Lys	His	Leu 65	Gly	Ala	Arg	Lys	Asp 70	His	Pro	Gly	Leu	Lys 75
Ala	Lys	Glu	Asn	Asp 80	Glu	Asn	Cys	Gly	Pro 85	Thr	Thr	Thr	Val	Phe 90
Val	Gly	Asn	Ile	Ser 95	Glu	Lys	Ala	Ser	Asp 100	Met	Leu	Ile	Arg	Gln 105
Leu	Leu	Ala	Lys	Cys 110	Gly	Leu	Val	Leu	Ser 115	Trp	Lys	Arg	Val	Gln 120
Gly	Ala	Ser	Gly	Lys 125	Leu	Gln	Ala	Phe	Gly 130	Phe	Cys	Glu	Tyr	Lys 135
Glu	Pro	Glu	Ser	Thr 140	Leu	Arg	Ala	Leu	Arg 145	Leu	Leu	His	Asp	Leu 150
Gln	Ile	Gly	Glu		Lys	Leu	Leu	Val	Lys 160	Val	Asp	Ala	Lys	Thr 165
Lys	Ala	Gln	Leu		Glu	Trp	Lys	Ala	Lys 175	Lys	Lys	Ala	Ser	Asn 180
Gly	Asn	Ala	Arg		Glu	Thr	Val	Thr	Asn	Asp	Asp	Glu	Glu	Ala

				185					190					195
Leu	Asp	Glu	Glu	Thr 200	Lys	Arg	Arg	Asp	Gln 205	Met	Ile	Lys	Gly	Ala 210
Ile	Glu	Val	Leu	Ile	Arg	Glu	Tyr	Ser	Ser 220	Glu	Leu	Asn	Ala	Pro
Ser	Gln	Glu	Ser	215 Asp	Ser	His	Pro	Arg		Lys	Lys	Lys	Glu	Lys
				230					235					240
Lys	Glu	Asp	lle	Phe 245	GŢĀ	Arg	rne	GIN	250	АТа	нIS			

## (2) INFORMATION FOR SEQ ID NO: 76:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 523 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single (D) TOPOLOGY: linear

## (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LNODNOT05 (B) CLONE: 3125156

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 76:

Met	Gly	Pro	Gln	Ala 5	Ala	Pro	Leu	Thr	Ile 10	Arg	Gly	Pro	Ser	Ser 15
				20	Pro				25					30
				35	Gly				40					45
				50	Leu				55					60
				65	Glu				70					75
_				80	Leu				85					Ala 90
				95	Gln				100					Pro 105
				110	Arg				115					120
				125	Leu				130					135
	-			140	Asp				145					150
				155	Arg				160					165
				170	Asp				175					Gly 180
Lys	Asp	Asp	Tyr	Ile 185	Asn	Ala	Ser	Cys	Val 190	Glu	Gly	Leu	Ser	Pro 195
_	_			200	Val				205					210
				215	Leu				220					225
				230	Ser				235					240
Ala	Arg	Tyr	Phe	Pro 245	Thr	Glu	Arg	Gly	Gln 250	Pro	Met	Val	His	Gly 255

```
Ala Leu Ser Leu Ala Leu Ser Ser Val Arg Ser Thr Glu Thr His
                                    265
                260
Val Glu Arg Val Leu Ser Leu Gln Phe Arg Asp Gln Ser Leu Lys
                                    280
                275
Arg Ser Leu Val His Leu His Phe Pro Thr Trp Pro Glu Leu Gly
                                    295
                290
Leu Pro Asp Ser Pro Ser Asn Leu Leu Arg Phe Ile Gln Glu Val
                                    310
                305
His Ala His Tyr Leu His Gln Arg Pro Leu His Thr Pro Ile Ile
                                     325
                                                         330
                320
Val His Cys Ser Ser Gly Val Gly Arg Thr Gly Ala Phe Ala Leu
                                     340
                335
Leu Tyr Ala Ala Val Glu Glu Val Glu Ala Gly Asn Gly Ile Pro
                                    355
                350
Glu Leu Pro Gln Leu Val Arg Arg Met Arg Gln Gln Arg Lys His
                                    370
                365
Met Leu Gln Glu Lys Leu His Leu Arg Phe Cys Tyr Glu Ala Val
                                    385
                380
Val Arg His Val Glu Gln Val Leu Gln Arg His Gly Val Pro Pro
                                     400
                395
Pro Cys Lys Pro Leu Ala Ser Ala Ser Ile Ser Gln Lys Asn His
                                    415
                410
Leu Pro Gln Asp Ser Gln Asp Leu Val Leu Gly Gly Asp Val Pro
                                                         435
                425
                                    430
Ile Ser Ser Ile Gln Ala Thr Ile Ala Lys Leu Ser Ile Arg Pro
                                    445
                440
Pro Gly Gly Leu Glu Ser Pro Val Ala Ser Leu Pro Gly Pro Ala
                                     460
                455
Glu Pro Pro Gly Leu Pro Pro Ala Ser Leu Pro Glu Ser Thr Pro
                                     475
                 470
Ile Pro Ser Ser Ser Gln Thr Pro Phe Pro Pro His Tyr Leu Arg
                 485
                                     490
Leu Pro Ser Leu Arg Arg Ser Arg Gln Cys Leu Lys Pro Pro Ala
                500
                                     505
                                                         510
Arg Gly Pro Pro Pro Pro Pro Trp Asn Cys Trp Pro Pro
                 515
                                     520
```

### (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 621 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGTUT12
- (B) CLONE: 3129120

### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 77:

Met Gly Leu Leu Ser Asp Pro Val Arg Arg Arg Ala Leu Ala Arg 10 15 Leu Val Leu Arg Leu Asn Ala Pro Leu Cys Val Leu Ser Tyr Val 20 25 30 Ala Gly Ile Ala Trp Phe Leu Ala Leu Val Phe Pro Pro Leu Thr 35 40 45 Gln Arg Thr Tyr Met Ser Glu Asn Ala Met Gly Ser Thr Met Val 50 60

77:

Glu	Glu	Gln	Phe		Gly	Gly	Asp	Arg	Ala 70	Arg	Ala	Phe	Ala	Arg 75
Asp	Phe	Ala	Ala	65 His 80	Arg	Lys	Lys	Ser		Ala	Leu	Pro	Val	
Trp	Leu	Glu	Arg	Thr 95	Met	Arg	Ser	Val		Leu	Glu	Val	Tyr	Thr 105
				Arg 110					115				His	120
				Ser 125					130				Arg	133
				140					145				Pro	150
				155					160				Leu -	102
				Phe 170					175					Ile 180
				185					190				Ala	195
				200					205				Ser	210
				215					220				Ala	225
				230					235				Val	Z40
				245					250				Leu	255
				260					265				Gln Leu	270
				275					280					283
				290					295				Ser	300
				305					310				Val	312
				320					325				Lys	330
				335	)				340				Arg	345
				350	)				355				Leu	360
				365	· )				370	}			Tyr	3/5
				380	)				385	)				390
				395					400	)				Glu 405
				410	)				415	)				Gln 420
_				425	5				430	)				435
				440	)				445	5				His 450
				455	5				460	)				. Val 465
				470	)				475	5				His 480
				485	5				490	)				7 Trp 495
				500	0				505	5				1 Leu 510
Glz	y Cys	s Il	e Ala	a Lei	u Th:	r Ası	n Phe	e Se:	r Lei	ı Gly	y Phe	е тел	л Тег	ı Ala

				515					520					525
				530					535	Lys				Pro 540
-				545					550	Thr				555
Thr	Leu	Leu	Gly	Ser	Leu	Phe	Leu	Trp	Arg 565	Glu	Leu	Gln	Glu	Ala 570
Pro	Leu	Ser	Leu	Ala 575	Glu	Gly	Trp	Gln	Leu 580	Phe	Leu	Ala	Ala	Leu 585
Ala	Gln	Gly	Val	Leu 590	Glu	His	His	Thr	Tyr 595	Gly	Ala	Leu	Leu	Phe 600
Pro	Leu	Leu	Ser	Leu 605	Gly	Leu	Tyr	Pro	Cys 610	Trp	Leu	Leu	Phe	Trp 615
Asn	Val	Leu	Phe	Trp 620	Lys									

#### (2) INFORMATION FOR SEQ ID NO: 78:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2347 base pairs

  - (B) TYPE: nucleic acid (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: HEARNOT01
  (B) CLONE: 305841
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 78:

CCCTCGAGAA	GATGGCGGCG	ACTCTGGGAC	CCCTTGGGTC	GTGGCAGCAG	TGGCGGCGAT	60
GTTTGTCGGC	TCGGGATGGG	TCCAGGATGT	TACTCCTTCT	TCTTTTGTTG	GGGTCTGGGC	120
AGGGGCCACA	GCAAGTCGGG	GCGGGTCAAA	CGTTCGAGTA	CTTGAAACGG	GAGCACTCGC	180
TGTCGAAGCC	CTACCAGGGT	GTGGGCACAG	GCAGTTCCTC	ACTGTGGAAT	CTGATGGGCA	240
ATGCCATGGT	GATGACCCAG	TATATCCGCC	TTACCCCAGA	TATGCAAAGT	AAACAGGGTG	300
CCTTGTGGAA	CCGGGTGCCA	TGTTTCCTGA	GAGACTGGGA	GTTGCAGGTG	CACTTCAAAA	360
TCCATGGACA	AGGAAAGAAG	AATCTGCATG	GGGATGGCTT	GGCAATCTGG	TACACAAAGG	420
ATCGGATGCA	GCCAGGGCCT	GTGTTTGGAA	ACATGGACAA	ATTTGTGGGG	CTGGGAGTAT	480
TTGTAGACAC	CTACCCCAAT	GAGGAGAAGC	AGCAAGAGCG	GGTATTCCCC	TACATCTCAG	540
CCATGGTGAA	CAACGGCTCC	CTCAGCTATG	ATCATGAGCG	GGATGGGCGG	CCTACAGAGC	600
TGGGAGGCTG	CACAGCCATT	GTCCGCAATC	TTCATTACGA	CACCTTCCTG	GTGATTCGCT	660
ACGTCAAGAG	GCATTTGACG	ATAATGATGG	ATATTGATGG	CAAGCATGAG	TGGAGGGACT	720
GCATTGAAGT	GCCCGGAGTC	CGCCTGCCCC	GCGGCTACTA	CTTCGGCACC	TCCTCCATCA	780
CTGGGGATCT	CTCAGATAAT	CATGATGTCA	TTTCCTTGAA	GTTGTTTGAA	CTGACAGTGG	840



AGAGAACCCC	AGAAGAGGAA	AAGCTCCATC	GAGATGTGTT	CTTGCCCTCA	GTGGACAATA	900
TGAAGCTGCC	TGAGATGACA	GCTCCACTGC	CGCCCTGAG	TGGCCTGGCC	CTCTTCCTCA	960
TCGTCTTTTT	CTCCCTGGTG	TTTTCTGTAT	TTGCCATAGT	CATTGGTATC	ATACTCTACA	1020
ACAAATGGCA	GGAACAGAGC	CGAAAGCGCT	TCTACTGAGC	CCTCCTGCTG	CCACCACTTT	1080
TGTGACTGTC	ACCCATGAGG	TATGGAAGGA	GCAGGCACTG	GCCTGAGCAT	GCAGCCTGGA	1140
GAGTGTTCTT	GTCTCTAGCA	GCTGGTTGGG	GACTATATTC	TGTCACTGGA	GTTTTGAATG	1200
CAGGGACCCC	GCATTCCCAT	GGTTGTGCAT	GGGGACATCT	AACTCTGGTC	TGGGAAGCCA	1260
CCCACCCCAG	GGCAATGCTG	CTGTGATGTG	CCTTTCCCTG	CAGTCCTTCC	ATGTGGGAGC	1320
AGAGGTGTGA	AGAGAATTTA	CGTGGTTGTG	ATGCCAAAAT	CACAGAACAG	AATTTCATAG	1380
CCCAGGCTGC	CGTGTTGTTT	GACTCAGAAG	GCCCTTCTAC	TTCAGTTTTG	AATCCACAAA	1440
GAATTAAAAA	CTGGTAACAC	CACAGGCTTT	CTGACCATCC	ATTCGTTGGG	TTTTGCATTT	1500
GACCCAACCC	TCTGCCTACC	TGAGGAGCTT	TCTTTGGAAA	CCAGGATGGA	AACTTCTTCC	1560
CTGCCTTACC	TTCCTTTCAC	TCCATTCATT	GTCCTCTCTG	TGTGCAACCT	GAGCTGGGAA	1620
AGGCATTTGG	ATGCCTCTCT	GTTGGGGCCT	GGGGCTGCAG	AACACACCTG	CGTTTCACTG	1680
GCCTTCATTA	GGTGGCCCTA	GGGAGATGGC	TTTCTGCTTT	GGATCACTGT	TCCCTAGCAT	1740
GGGTCTTGGG	TCTATTGGCA	TGTCCATGGC	CTTCCCAATC	AAGTCTCTTC	AGGCCCTCAG	1800
TGAAGTTTGG	CTAAAGGTTG	GTGTAAAAAT	CAAGAGAAGC	CTGGAAGACA	TCATGGATGC	1860
CATGGATTAG	CTGTGCAACT	GACCAGCTCC	AGGTTTGATC	AAACCAAAAG	CAACATTTGT	1920
CATGTGGTCT	GACCATGTGG	AGATGTTTCT	GGACTTGCTA	GAGCCTGCTT	AGCTGCATGT	1980
TTTGTAGTTA	CGATTTTTGG	AATCCCACTT	TGAGTGCTGA	AAGTGTAAGG	AAGCTTTCTT	2040
CTTACACCTT	GGGCTTGGAT	ATTGCCCAGA	GAAGAAATTT	GGCTTTTTTT	TTCTTAATGG	2100
ACAAGAGACA	GTTGCTGTTC	TCATGTTCCA	AGTCTGAGAG	CAACAGACCC	TCATCATCTG	2160
TGCCTGGAAG	AGTTCACTGT	CATTGAGCAG	CACAGCCTGA	GTGCTGGCCT	CTGTCAACCC	2220
TTATTCCACT	GCCTTATTTG	ACAAGGGGTT	ACATGCTGCT	CACCTTACTG	CCCTGGGATT	2280
AAATCAGTTA	CAGGCCAGAG	TCTCCTTGGA	GGGCCTGGAA	CTCTGAGTCC	TCCTATGAAC	2340
CTCTGTA						2347

## (2) INFORMATION FOR SEQ ID NO: 79:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1529 base pairs (B) TYPE: nucleic acid

- (C) STRANDEDNESS: single
  (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: EOSIHETO2 (B) CLONE: 322866
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 79:

(2117)			~			
CCCACGCGTC	CGCCAGCCTT	GTCTCGGCCA	CCTCAAGGAT	AATCACTAAA	TTCTGCCGAA	60
AGGACTGAGG	AACGGTGCCT	GGAAAAGGGC	AAGAATATCA	CGGCATGGGC	ATGAGTAGCT	120
TGAAACTGCT	GAAGTATGTC	CTGTTTTTCT	TCAACTTGCT	CTTTTGGATC	TGTGGCTGCT	180
GCATTTTGGG	CTTTGGGATC	TACCTGCTGA	TCCACAACAA	CTTCGGAGTG	CTCTTCCATA	240
ACCTGCCCTC	CCTCACGCTG	GGCAATGTGT	TTGTCATCGT	GGGCTCTATT	ATCATGGTAG	300
TTGCCTTCCT	GGGCTGCATG	GGCTCTATCA	AGGAAAACAA	GTGTCTGCTT	ATGTCGTTCT	360
TCATCCTGCT	GCTGATTATC	CTCCTTGCTG	AGGTGACCTT	GGCCATCCTG	CTCTTTGTAT	420
ATGAACAGAA	GCTGAATGAG	TATGTGGCTA	AGGGTCTGAC	CGACAGCATC	CACCGTTACC	480
ACTCAGACAA	TAGCACCAAG	GCAGCGTGGG	ACTCCATCCA	GTCATTTCTG	CAGTGTTGTG	540
GTATAAATGG	CACGAGTGAT	TTGGACAGTG	GCTCACCAGC	ATCTTGCCCC	TCAGATCGAA	600
AAGTGGAGGG	GTGCTATGCG	AAAGAAGACT	TTGGTTTCAT	TCAATTTCCT	GTATATCGGA	660
ATCATCACCA	TCTGTGTATG	TGTGATTGAG	GTGTTGGGGG	ATGTCCTTTG	CACTGACCCT	720
GAACTGCCAG	ATTGACAAAA	CCAGCCAGAC	CATAGGGCTA	TGATCTGCAG	TAGTTCTGTG	780
GTGAAGAGAC	TTGTTTCATC	TCCGGAAATG	CAAAACCATT	TATAGCATGA	AGCCCTACAT	840
GATCACTGCA	GGATGATCCT	CCTCCCATCC	TTTCCCTTTT	TAGGTCCCTG	TCTTATACAA	900
CCAGAGAAGT	GGGTGTTGGC	CAGGCACATC	CCATCTCAGG	CAGCAAGACA	ATCTTTCACT	960
CACTGACGGC	AGCAGCCATG	TCTCTCAAAG	TGGTGAAACT	AATATCTGAG	CATCTTTTAG	1020
ACAAGAGAGG	CAAAGACAAA	. CTGGATTTAA	TGGCCCAACA	TCAAAGGGTG	AACCCAGGAT	1080
ATGAATTTTT	GCATCTTCCC	ATTGTCGAAT	TAGTCTCCAG	CCTCTAAATA	ATGCCCAGTC	1140
TTCTCCCCAA	AGTCAAGCAA	. GAGACTAGTT	GAAGGGAGTT	CTGGGGCCAG	GCTCACTGGA	1200
CCATTGTCAC	AACCCTCTGT	TTCTCTTTGA	CTAAGTGCCC	TGGCTACAGG	AATTACACAG	1260
TTCTCTTTCT	CCAAAGGGCA	AGATCTCATT	TCAATTTCTT	' TATTAGAGGG	G CCTTATTGAT	1320
GTGTTCTAAG	TCTTTCCAGA	AAAAAACTAT	CCAGTGATT	ATATCCTGAT	TTCAACCAGT	1380
CACTTAGCTG	ATAATCACAG	TAAGAAGACI	TCTGGTATTA	TCTCTCTATC	C AGATAAGATT	1440
TTGTTAATGT	CACTATTTAC	C TCTTCAATAA	ATAAAACAGI	TTATTATCTC	C AAAAAAAAA	1500
AAAAAAAA	AAAAAAAA	AAAAAAAA				1529

# (2) INFORMATION FOR SEQ ID NO:

- (i) SEQUENCE CHARACTERISTICS:
  - $(\bar{A})$  LENGTH: 4387 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
    (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BEPINOT01
- (B) CLONE: 546656
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 80 :

• •						
GCATCCCCGC	TTCCGGGTTA	GGCCGTTCCT	GCCCGCCCCC	TCCTCTCCTC	CCTTCGGACC	60
CATAGATCTC	AGGCTCGGCT	CCCCGCCCGC	CGCAGCCCAC	TGTTGACCCG	GCCCGTACTG	120
CGGCCCCGTG	GCCACCATGT	CCCTGCACGG	CAAACGGAAG	GAGATCTACA	AGTATGAAGC	180
GCCCTGGACA	GTCTACGCGA	TGAACTGGAG	TGTGCGGCCC	GATAAGCGCT	TTCGCTTGGC	240
GCTGGGCAGC	TTCGTGGAGG	AGTACAACAA	CAAGGTTCAG	CTTGTTGGTT	TAGATGAGGA	300
GAGTTCAGAG	TTTATTTGCA	GAAACACCTT	TGACCACCCA	TACCCCACCA	CAAAGCTCAT	360
GTGGATCCCT	GACACAAAAG	GCGTCTATCC	AGACCTACTG	GCAACAAGCG	GTGACTATCT	420
CCGTGTGTGG	AGGGTTGGTG	AAACAGAGAC	CAGGCTGGAG	TGTTTGCTAA	ACAATAATAA	480
GAACTCTGAT	TTCTGTGCTC	CCCTGACCTC	CTTTGACTGG	AATGAGGTGG	ATCCTTATCT	540
TTTAGGTACC	TCAAGCATTG	ATACGACATG	CACCATCTGG	GGGCTGGAGA	CAGGGCAGGT	600
GTTAGGGCGA	GTGAATCTCG	TGTCTGGCCA	CGTGAAGACC	CAGCTGATCG	CCCATGACAA	660
AGAGGTCTAT	GATATTGCAT	TTAGCCGGGC	CGGGGGTGGC	AGGGACATGT	TTGCCTCTGT	720
GGGTGCTGAT	GGCTCGGTGC	GGATGTTTGA	CCTCCGCCAT	CTAGAACACA	GCACCATCAT	780
TTACGAAGAC	CCACAGCATC	ACCCACTGCT	TCGCCTCTGC	TGGAACAAGC	AGGACCCTAA	840
CTACCTGGCC	ACCATGGCCA	TGGATGGAAT	GGAGGTGGTG	ATTCTAGATG	TCCGGGTTCC	900
CTGCACACCT	GTCGCCAGGT	TAAACAACCA	TCGAGCATGT	GTCAATGGCA	TTGCTTGGGC	960
CCCACATTCA	TCCTGCCACA	TCTGCACTGC	AGCGGATGAC	CACCAGGCTC	TCATCTGGGA	1020
CATCCAGCAA	ATGCCCCGAG	CCATTGAGGA	CCCTATCCTG	GCCTACACAG	CTGAAGGAGA	1080
GATCAACAAT	GTGCAGTGGG	CATCAACTCA	GCCCGACTGG	ATCGCCATCI	GCTACAACAA	1140
CTGCCTGGAG	ATACTCAGAG	TGTAGTGTTG	GTGGCGCTGT	' GCCCACGAGG	CAGGGGCTTT	1200
TGTATTTCCT	GCCTCTGCCC	CACCCCAAA	GTAAGAAGAA	ACATGTTTCC	AGTGGCCAGT	1260
ATGTCTTTCA	TTGCTTTGCA	CCCACTGTTA	CCAGAAGCTG	CTCTAGGAGT	TCCTGGCCAG	1320

PF-0459 US TCACCCCATC GCCCTCTGTG GCAGACTCAG TGCTGTGTGG CGCCTCCTCA GCCCAGGGCT 1380 GAGTTTTAAG ATTTTCTCTC CTTTCCTCTT CTCCTTTGGT TCCTCAATTA AAAAATGTGT 1440 GTATATTTGT TTGTCAGGCG TTGTGTTGAG GAGCAGTTCA CGCACTGGCT GTGTCTATTC 1500 CTCTGCCCAG GTGTCTCTGT TTGCTGCCCA AGGCAGCAGT TCATGTCTCG TCCATGTCCA 1560 TGTTCGTGTT AGCACTTACG TGGGAACAAA TACCAATTTG TCTTTTCTCC TAGTATCAGT 1620 GTGTTTAACA AATTTTAACT TTGTATATTT GTTATCTATC AGGCTAATTT TTTTATGAAA 1680 AGAATTTTAC TCTCCTGCTT CATTTCTTTG TCTTATAGTC CTCCCTCTTT GCACCTTCTT 1740 CTCTTCCCTC AGTGCCTGGA GCTGGTACTG GGCCCCTGGG CCCCATGAGC AGTTTGCCTT 1800 CTTGAGTCAC TGCCTGTGTA GTACATACCT GACCGGGAGT CCAAACCACC TTGGTGCTCT 1860 GAAGTCCACT GACTCATCAC ACCTTTCTTA GCCTGGCTCC TCTCAAGGGC ATTCTGGGCT 1920 TGTAAACAGA CATAGGAAGC CTCTGTTTAC CCTGAAGCAC CACTGTCCAG CCCATTGGTT 1980 CCCACTGGCA GCATGGTAGA GCTGAGAGAA ACAGGCTCTC AGGGTACCTG ACTTGAGGGG 2040 AATCGTTTCA TGAAGCTGAA CTTCAAGCAT ATTTCCAGTA CATTCTTTCA GAGTCTGTTT 2100 TTCCATCCAA ATATAAGCCC CAGGCCATTC CACTTAGTGT CTTTTCAATG ATAGGCAAGA 2160 ATGATATCTG AGTTGAACTT CGGTGCTTCT GTTGTTTGAG TTTACTGTGC CTGGTGGTAT 2220 ATTGGGCATT CTTTGGATTG AGTGTTCTGA GGTGAGAGAG TCTTCCCGAG GCATCCTGTC 2280 TGTGCTTCCA ACCCTGAACA AGACCTTACA TGAGAGATGG ACTGATGGAC TGCGGCAATC 2340 CTGGGCTGTC AAGTGGATAG ATAGTTAAAA AGCATTATAC TGTGGGTAAT GAAAAGGGAG 2400 GAAAAAAAA GAAGGAAAAG GAATTATAGA CCCCCAGGGT CAGCCAGTTA AGAGCTCTAC 2460 CCACACCTGT CAACCCCTCT CTCCCCCAGT TTAGGTTCTG AGCAGTATTG GACTTGTAGC 2520 CTGCAGTTGT CTTTTGACTT GCAGGCCGCA GGTGTCTTTC TGTTATGTGA ATGAGTTCCA 2580 TGGAGGGGCA TATGTGTGAT TCCACCGTTA GATGAGCCCT TGGGGCAGGC AGTTTGGGAT 2640 GTGCTCTTGG GGGAAAGTTG GCTGTTTCCT TGCGCTCTGC TCCTACCCGA AGGTTTTTAA 2700 GTCCCTCTGA ATTGCTCATC TGAGATTAGT AGAGTAGCAG GCCTGAAGGA TGATGGTTTT 2760 GTCCTCTTTG GTTCTCACCT GCTTGAGAAG TAAAACAGTA ACTTTGTTCT TCTGGGCCCT 2820 TAAGCTTTTT TGGTTAAGTC TTCCTTTTCA GAAGTAGATG TCATTATATG CCAAAAGTCT 2880 AGCTCTTTGC TTTACCATAC AGGGACCTGT CCCAAAGAAA AAGGCTCTTT TTTTAGCCAG 2940 CATATTTCCC CTTCTACCCT TTTACTTTGT TGTTCTGATT TTAGGACTCT GGCTGGCCAT 3000 GTGCTTGTGG TTGCCTCTCC TGCATTTGCC ACTGGATTTG CACTGCATCG TTTGGAGATA 3060 CAAAGCGAGC AGTTCTTGGT CAGAACCCTC CTCTGCTTTT CATTGTGTTT GATAATGGTT 3120

ACTGGGTCCT TCTCTCAAGG GTAGCAAGGC CAAGCTGATG GCTGCTTGTT TAGGAGGCCA 3180

TCAGTTCCTT	CCTGTGGAGA	AGGGTCTGAA	ATGGAAGTCA	GTGGTAGAAG	GGGCTGGTCT	3240
GCTGGGCAGG	GCTTACATCC	ACTGAGTTCT	AAGATTCCTT	TCCTGATCTG	CACCTACGCC	3300
TGGTCTGTAT	GGTGGAATTT	GTCAGCTGGA	ACTCAGAAAC	AACAACTTGA	AAAAAAAATA	3360
ATAATTAGAA	CATATTTGCA	TAAGATAGCT	ATTTACTCTG	GAAACCAACA	ACTTTTGAGA	3420
TTTCCCTTGC	CCTGTGGACG	CCCAGCTCCT	GTCATCCTTC	CTTAGGTCCT	GCAGTACAGT	3480
CTTCCCCTGA	ATGCCACCGG	GGACCCAGGG	GGACTCCACC	CCCCTAAGCA	AGCACACACA	3540
TACTCACAGT	TGATGAGTTG	CTGGTCTTTG	AGTCCCAGCT	CTCTTACCCT	CCCTTTACTC	3600
CACCAGCCCG	ACGACCCATG	ACTGAGGAGG	GGATTTCTAC	AGTCTCAGGA	TTTAGAAAGT	3660
CTGTAAGCCA	TCCATGCTCC	AGAAAGCACC	GATCTGTTGT	AGTTGCAAAA	ACAACTCTGT	3720
AATTTGTTGA	GGTTCTCAAA	CTGACAGCCA	GCGAGACTGG	GTGGGAGGCC	CTGGATCTGT	3780
TCTCCCTGAC	TGCGGGAGGA	GCAGCCACTA	GGACTTTAGC	AGGAAGCCCA	CATGGAGGCT	3840
CCGCCAGGCT	GTGGCCCAGC	TGGTGATGGC	CCTTTTGCTC	CTGGCAGCCT	GAGGCACAGC	3900
TGCCTGTATT	GTCCTCATCT	GTTCTGACTG	AAGGATGGAG	GTGCTGAATA	AATTAGGCCT	3960
CAGGCCTCTA	CCACCAGAGA	GCTGGAGAAT	GGGTCCACGT	CATTCAAGGA	CCTGAATTTT	4020
TTATGCTCAG	GAGCATTGGA	ATCCTCTTCT	TCCAGGGAGG	AATTAGCCTG	CAAGGTTAGG	4080
ACTTGAAGAG	GGAAGGTATT	TAATAACTGG	GCGAGGATGG	GTGTGGTGGC	TCACACCTGT	4140
AATCCCAGCA	TTTTGGGAGG	CTGAGGTGGC	CAGATCCCAA	GGTCAGAAGA	. TCGAGACCAT	4200
CCTGGCTAAC	ATGGTGAAAC	CCCATCTCTA	CTAAAAATAC	TAAAAAAA	TAGCCGGGGG	4260
TGGTGGCGGG	TACCTGTAGT	CCTAGCTACT	TGGGAGGCTG	AGGCAGGAGA	ATGGCGTGAA	4320
CCTGGGAGGT	GGAGCTTGCA	GTGAGCCAAG	ATCGTCCACT	CACTGCAGCC	: TGGCGACAGA	4380
GCAAGCG						4387

#### (2) INFORMATION FOR SEQ ID NO: 81:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2117 base pairs

  - (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: SYNORAT03
  - (B) CLONE: 693453
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 81 :

GCCTGAGCGG GAAGCATTGG CGTCCGAGCG ACTTCTAGGA GCCTGGGGTT CGGCGCTATG 60



GAGGAGCTCG	ATGGCGAGCC	AACAGTCACT	TTGATTCCAG	GCGTGAATTC	CAAGAAGAAC	120
CAAATGTATT	TTGACTGGGG	TCCAGGGGAG	ATGCTGGTAT	GTGAAACCTC	CTTCAACAAA	180
AAAGAAAAAT	CAGAGATGGT	GCCAAGTTGC	CCCTTTATCT	ATATCATCCG	TAAGGATGTA	240
GATGTTTACT	CTCAAATCTT	GAGAAAACTC	TTCAATGAAT	CCCATGGAAT	CTTTCTGGGC	300
CTCCAGAGAA	TTGACGAAGA	GTTGACTGGA	AAATCCAGAA	AATCTCAATT	GGTTCGAGTG	360
AGTAAAAACT	ACCGATCAGT	CATCAGAGCA	TGTATGGAGG	AAATGCACCA	GGTTGCAATT	420
GCTGCTAAAG	ATCCAGCCAA	TGGCCGCCAG	TTCAGCAGCC	AGGTCTCCAT	TTTGTCAGCA	480
ATGGAGCTCA	TCTGGAACCT	GTGTGAGATT	CTTTTTATTG	AAGTGGCCCC	AGCTGGCCCT	540
CTCCTCCTCC	ATCTCCTTGA	CTGGGTCCGG	CTCCATGTGT	GCGAGGTGGA	CAGTTTGTCG	600
GCAGATGTTC	TGGGCAGTGA	GAATCCAAGC	AAACATGACA	GCTTCTGGAA	CTTGGTGACC	660
ATCTTGGTGC	TGCAGGGCCG	GCTGGATGAG	GCCCGACAGA	TGCTCTCCAA	GGAAGCCGAT	720
GCCAGCCCCG	CCTCTGCAGG	CATATGCCGA	ATCATGGGGG	ACCTGATGAG	GACAATGCCC	780
ATTCTTAGTC	CTGGGAACAC	CCAGACACTG	ACAGAGCTGG	AGCTGAAGTG	GCAGCACTGG	840
CACGAGGAAT	GTGAGCGGTA	CCTCCAGGAC	AGCACATTCG	CCACCAGCCC	TCACCTGGAG	900
TCTCTCTTGA	AGATTATGCT	GGGAGACGAA	GCTGCCTTGT	TAGAGCAGAA	GGAACTTCTG	960
AGTAATTGGT	ATCATTTCCT	AGTGACTCGG	CTCTTGTACT	CCAATCCCAC	AGTAAAACCC	1020
ATTGATCTGC	ACTACTATGC	CCAGTCCAGC	CTGGACCTGT	TTCTGGGAGG	TGAGAGCAGC	1080
CCAGAACCCC	TGGACAACAT	CTTGTTGGCA	GCCTTTGAGT	TTGACATCCA	TCAAGTAATC	1140
AAAGAGTGCA	GCATCGCCCT	GAGCAACTGG	TGGTTTGTGG	CCCACCTGAC	AGACCTGCTG	1200
GACCACTGCA	AGCTCCTCCA	GTCACACAAC	CTCTATTTCG	GTTCCAACAT	GAGAGAGTTC	1260
CTCCTGCTGG	AGTACGCCTC	GGGACTGTTT	GCTCATCCCA	GCCTGTGGCA	GCTGGGGGTC	1320
GATTACTTTG	ATTACTGCCC	CGAGCTGGGC	CGAGTCTCCC	TGGAGCTGCA	CATTGAGCGG	1380
ATACCTCTGA	ACACCGAGCA	GAAAGCCCTG	AAGGTGCTGC	GGATCTGTGA	GCAGCGGCAG	1440
ATGAC <b>T</b> GAAC	AAGTTCGCAG	CATTTGTAAG	ATCTTAGCCA	TGAAAGCCGT	CCGCAACAAT	1500
CGCCTGGGTT	CTGCCCTCTC	TTGGAGCATC	CGTGCTAAGG	ATGCCGCCTT	TGCCACGCTC	1560
GTGTCAGACA	GGTTCCTCAG	GGATTACTGT	GAGCGAGGCT	GCTTTTCTGA	TTTGGATCTC	1620
ATTGACAACC	TGGGGCCAGC	CATGATGCTC	AGTGACCGAC	TGACATTCCT	GGGAAAGTAT	1680
CGCGAGTTCC	ACCGTATGTA	CGGGGAGAAG	CGTTTTGCCG	ACGCAGCTTC	TCTCCTTCTG	1740
TCCTTGATGA	. CGTCTCGGAT	TGCCCCTCGG	TCTTTCTGGA	TGACTCTGCT	GACAGATGCC	1800
TTGCCCCTTT	TGGAACAGAA	ACAGGTGATT	TTCTCAGCAG	AACAGACTTA	. TGAGTTGATG	1860
CGGTGTCTGG	AGGACTTGAC	GTCAAGAAGA	CCTGTGCATG	GAGAATCTGA	. TACCGAGCAG	1920

### (2) INFORMATION FOR SEQ ID NO: 82:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 846 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: BRAITUT03
  - (B) CLONE: 866885
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 82:
- GGCGGGCGGA GTCTGCAGGA TGGCACCGGA CCCCTGGTTC TCCACATACG ATTCTACTTG 60 TCAAATTGCC CAAGAAATTG CTGAGAAAAT TCAACAACGA AATCAATATG AAAGAAAAGG 120 TGAAAAGGCA CCAAAGCTTA CCGTGACAAT CAGAGCTTTG TTGCAGAACC TGAAGGAAAA 180 GATCGCCCTT TTGAAGGACT TATTGCTAAG AGCTGTGTCA ACACATCAGA TAACACAGCT 240 TGAAGGGGAC CGAAGACAGA ACCTCTTGGA TGATCTTGTA ACTCGAGAGA GACTACTTCT 300 GGCATCCTTT AAGAATGAGG GTGCCGAACC AGATCTAATC AGGTCCAGCC TGATGAGTGA 360 AGAGGCTAAG CGAGGAGCAC CCAACCCTTG GCTCTTTGAG GAGCCAGAGG AGACCAGAGG 420 CTTGGGTTTT GATGAAATCC GGCAACAGCA GCAGAAAATT ATCCAAGAAC AGGATGCAGG 480 CCTTGATGCC CTTTCCTCTA TCATAAGTCG CCAAAAACAA ATGGGGCAGG AAATTGGGAA 540 TGAATTGGAT GAACAAAATG AGATAATTGA CGACCTTGCC AACCTAGTGG AGAACACAGA 600 TGAAAAACTT CGCAATGAAA CCAGGCGGGT AAACATGGTG GACAGAAAGT CAGCCTCTTG 660 TGGGATGATC ATGGTGATTT TACTGCTGCT TGTGGCTATC GTGGTTGTTG CAGTCTGGCC 720 GACCAACTGA TGGCAGTAAA GAGACCACCA GCAGTGACAC CTGGCAATGA CAGATGCAAG 780 CCCAACACCC TTTTGGTACG CAAAACCTGC TCTCAATAAA TTCCCCCAAA GCTCTGAAAA 840 846 AAAAAA
- (2) INFORMATION FOR SEQ ID NO: 83:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1011 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LUNGNOT03
  - (B) CLONE: 1242271
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 83:
- GAAAGAGATA ACTGGAAGTT CCTTGATTCA GAAAACAGAT TCAGATGAAG AAGTTGCAAT GCTGTTGGAC ACAGTCCAGA AAGTATTTCA GAAAATGTTG GAATGTATTG CACGGAGCTT 120 CAGGAAGCAG CCGGAAGAAG GCCTGCGGCT GCTTTATTCT GTTCAGAGGC CTCTTCATGA 180 GTTCATTACT GCTGTTCAGT CTCGGCACAC AGACACCCCT GTGCACCGGG GTGTACTTTC 240 TACTCTGATC GCTGGGCCTG TGGTTGAGAT AAGTCACCAG CTACGGAAGG TTTCTGACGT 300 AGAAGAGCTT ACCCCTCCAG AGCATCTTTC TGATCTTCCA CCATTTTCAA GGTGTTTAAT 360 AGGAATAATA ATAAAGTCTT CGAATGTGGT CAGGTCATTT TTGGATGAAT TAAAGGCATG 420 TGTGGCTTCT AATGATATTG AAGGCATTGT GTGCCTCACG GCTGCTGTGC ATATTATCCT 480 GGTTATTAAT GCAGGTAAAC ATAAAAGCTC AAAAGTGAGG GAGGTTGCAG CCACTGTTCA 540 CAGAAAACTA AAGACATTCA TGGAAATTAC TTTGGAAGAG GATAGCATTG AAAGATTTCT 600 CTATGAATCA TCATCAAGAA CTCTGGGAGA ACTTTTGAAT TCATAACCAA GCCAACATCT 660 CCAGACATGT AAAAATAGGG AAAAGTGATT CAAATTGAAA TGCCTGTGTA TTTTCCTATT 720 GTTTTTAATG TTAATAACCC ATATAATAGG GAAAGGGTGG GATTTTTTTG TGGGAATGTG 780 GGAAGGTGGG GGTTATGGAG GAGATAACTC AAAACTTCTT CAATTTTGCC TAGTGCCTGC 840 GTAAATAATA TATTTAATAT AAAGGACTCC AGGTATGAAT GGTGTAGAAA TCCATGATTC 900 CAAGAAAAA CACTTTTCTA GCAAACCTGG TTGTTTTTAA AATGACTTTT ATATATGTAA 960 TATTGCTTGG AAACTATGAG TAATAAAGCA ATGACAACAT CAAAAAAAAA A
- (2) INFORMATION FOR SEQ ID NO: 84:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 2478 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: LUNGFET03
    - (B) CLONE: 1255027

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 84:

CCCACGCGTC CGCCCACGCG TCCGCAGCGC TGTGTTTGCG AGCGGGAGCG AGGGGCGCCG 60 GCTGGGGTGT GTGCTCCTGA GCTCTTCAGA AACCAGGCTG CTTTCAGGAA CATTGCTGTG 120 GATTCCCAGC TTTCAGACAA CACATGACTA AGACAGATGA GACCACTCTA GTTGCCTCAT 180 GGGAAACTCG GGAAAAGACT GCAAAAACAA CATTGTTTCT CCCTTTGGAA TTCTGGAGTT 240 ATAAGGCAGA GGTCCCCCAT CTTCCCGAAC TGGCCTATTC CGCTAGAAGC AAGATGGCTG 300 AACTCAATAC TCATGTGAAT GTCAAGGAAA AGATCTATGC AGTTAGATCA GTTGTTCCCA 360 ACAAAAGCAA TAATGAAATA GTCCTGGTGC TCCAACAGTT TGATTTTAAT GTGGATAAAG 420 CCGTGCAAGC CTTTGTGGAT GGCAGTGCAA TTCAAGTTCT AAAAGAATGG AATATGACAG 480 GCAAAAAGAA GAACAATAAA AGAAAAAGAA GCAAGTCCAA GCAGCATCAA GGCAACAAAG 540 ATGCTAAAGA CAAGGTGGAG AGGCCTGAGG CAGGGCCCCT GCAGCCGCAG CCACCACAGA 600 TTCAAAACGG CCCCATGAAT GGCTGCGAGA AGGACAGCTC GTCCACAGAT TCTGCTAACG 660 AAAAACCAGC CCTTATCCCT CGTGAGAAAA AGATCTCGAT ACTTGAGGAA CCTTCAAAGG 720 CACTTCGTGG GGTCACAGAA GGCAACAGAC TACTGCAACA GAAACTATCC TTAGATGGGA 780 ACCCCAAACC TATACATGGA ACAACAGAGA GGTCAGATGG CCTACAGTGG TCAGCTGAGC 840 AGCCTTGTAA CCCAAGCAAG CCTAAGGCAA AAACATCTCC TGTTAAGTCC AATACCCCTG 900 CAGCTCATCT TGAAATAAAG CCAGATGAGT TGGCAAAGAA AAGAGGCCCA AATATTGAGA 960 AATCAGTGAA GGATTTGCAA CGCTGCACCG TTTCTCTAAC TAGATATCGC GTCATGATTA 1020 AGGAAGAAGT GGATAGTTCC GTGAAGAAGA TCAAAGCTGC CTTTGCTGAA TTACACAACT 1080 GCATCATTGA CAAAGAAGTT TCATTAATGG CAGAAATGGA TAAAGTTAAA GAAGAAGCCA 1140 TGGAAATCCT GACTGCTCGT CAGAAGAAAG CAGAAGAACT AAAGAGACTC ACTGACCTTG 1200 CCAGTCAGAT GGCAGAGATG CAGCTGGCCG AACTCAGGGC AGAAATTAAG CACTTTGTCA 1260 GCGAGCGTAA ATATGACGAG GAGCTCGGGA AAGCTGCCCG GTTTTCCTGT GACATCGAAC 1320 AGCTGAAGGC CCAAATCATG CTCTGCGGAG AAATTACACA TCCAAAGAAC AACTATTCCT 1380 CAAGAACTCC CTGCAGCTCC CTGCTGCCTC TGCTGAATGC GCACGCAGCA ACCTCTGGGA 1440 AACAGAGTAA CTTTTCCCGA AAATCATCCA CTCACAATAA GCCCTCTGAA GGCAAAGCGG 1500 CAAACCCCAA AATGGTGAGC AGTCTCCCCA GCACCGCCGA CCCCTCTCAC CAGACCATGC 1560 CGGCCAACAA GCAGAATGGA TCTTCTAACC AAAGACGGAG ATTTAATCCA CAGTATCATA 1620 ACAACAGGCT AAATGGGCCT GCCAAGTCGC AGGGCAGTGG GAATGAAGCC GAGCCACTGG 1680 GAAAGGGCAA CAGCCGCCAC GAACACAGAA GACAGCCGCA CAACGGCTTC CGGCCCAAAA 1740 ACAAAGGCGG TGCCAAAAAT CAAGAGGCTT CCTTGGGGAT GAAGACCCCC GAGGCCCCGG 1800

CCCATTCTGA AAAGCCCCGG CGAAGGCAGC ACGCTGCAGA CACCTCGGAG GCCAGGCCCT 1860

TCCGGGGTAG TGTCGGTAGG GTTTCACAGT GCAATCTCTG CCCCACGAGA ATAGAAGTTT 1920

CCACAGATGC AGCAGTTCTC TCAGTCCCGG CTGTGACGTT GGTGGCCTGA GCTAGGAGGA 1980

AAAAGAGCAG TTTTCACTCA GTTTTGGTTC CCTGCCCGAG GTGCTGACCC AATTCGCTGC 2040

CAAAAGAGTG TCAATCAGAA TATACAAAATC CCGTATGGTT GTGTCATCC CTCTTAATCA 2100

TTTTTACTAA TTCTAATAAT CAGCTCTAGC TTGCTTCATA ATTTTCATGG CTTTGCTTGA 2160

TCTGTTGATG CTTTCTCCA TCAAGACTTT GCAGCATTTT AGCCAGGCAG TATTTACTCA 2220

TTATTAGGAA AATCAAGATG TGGCTGAAGA TCAGAGGCTC AGTTAGCAAC CTGTGTTGTA 2340

ACAGTGATGT CAGTCCATTG ATTGTCTTTA GAGAGTTAAT GTTACAAAAA AGAATTCTTA 2340

ATAATCAGAC AAACATGATC TGCTGAGGAC ACATGCGCTT TTGTAGAATT TAACATCTGG 2400

TGCATTTGAA AAAAAAAA

### (2) INFORMATION FOR SEQ ID NO: 85:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1897 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: TESTTUT02
  - (B) CLONE: 1273453
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 85:
- AAGTGACATCTA GCACAAATTG AAGATGATAG AGCTGCGATG GTTATTTCTT GGCATCTGGC 60

  AAGTGACATG GACTGTGTAG TCACCCTAAC CACTGACGCT GCACGTCGTA TCTATGATGA 120

  AACCCAAGGT CGTCAGCAGG TGTTGCCCCT TGATTCTATT TACAAGAAGA CTCTTCCAGA 180

  TTGGAAAAGA TCTCTACCTC ATTTCCGAAA TGGAAAATTG TATTTTAAAC CCATTGGAGA 240

  TCCAGTCTTT GCTCGAGACT TGTTAACATT TCCAGATAAT GTAGAACATT GTGAAACAGT 300

  ATTTGGTATG CTGTTAGGAG ACACCATTAT TTTGGATAAT CTGGATGCGG CCAATCATTA 360

  TAGAAAAGAG GTTGTTAAAA TTACACACTG TCCTACACTG CTGACCAGAG ATGGAGATCG 420

  AATTCGAAGT AATGGAAAGT TTGGGGGCCT TCAGAATAAA GCTCCTCCAA TGGATAAACT 480

  TCGGGGAATG GTATTTGGAG CTCCAGTTCC AAAACAGTGT CTGATCTTAG GGGAACAAAT 540

  AGATCTTCTT CAGCAGTATC GTTCTGCTGT GTGCAAACTA GACAGTGTA ATAAGGATCT 600



TAACAGTCAA	TTAGAGTACC	TTCGCACTCC	GGATATGAGG	AAGAAAAAGC	AAGAACTTGA	660
TGAACATGAG	AAAAATCTCA	AACTAATAGA	GGAAAAACTA	GGTATGACTC	CCATACGTAA	720
GTGTAATGAC	TCATTGCGTC	ATTCACCAAA	GGTTGAGACG	ACAGATTGTC	CAGTTCCTCC	780
TAAAAGAATG	AGACGAGAAG	CTACAAGACA	AAATAGGATT	ATAACCAAAA	CAGATGTATG	840
AGAGGTGACA	GAGAGAAGAG	GCCATTGGTC	TCAGTAAGAA	TGCCCTGCTT	TCTGCATCTC	900
TGTTTCAGAA	GACCAAGAGG	GTGACTTACC	AGACTGAGTA	TTTCTGGGGA	CAATACAAGT	960
ACCTGGGCAT	GAATTTCCAT	TTCGATTCAG	ATGGGACTGG	AAACAACCAT	TCAATTTTAT	1020
GAATCTTACT	GGACATTATG	GATTTACTGG	AATTATTCCA	GACATTATGC	CCTTTGGTTG	1080
TCACTACCTT	GCAAATGTGT	AAGAGGAAAA	TGTGCTAATG	TGGCAGTGAC	TGTAAAACTG	1140
GCACATGGCA	TTTATTAATC	CTGAAGAAAA	GTACATGTAC	TATTTTCAG	TATAAATATA	1200
ATGAACATGT	CAGAACTATT	TCTTGAAAAC	CTTTTTATTA	CTTTTGCGTG	AATTTATTTA	1260
ACAAAGATGT	TTTGTCTTTT	GTGTAAGGGA	GGTTCTAGAG	GCTAGATGTT	TAATTGTAAA	1320
TATGTGAGGA	AACTCAATGC	AGAATTCAGG	ATAAAAATTT	TAAAAGCACA	GGTATTTGGG	1380
AATTGAAATG	TTAAGATACC	CAGAACAACA	TTAAATCAAT	GAGTGAACTT	GTGACAGTGG	1440
TAGCATTTCA	AATTTCAAAA	GACTTATCCT	GTGTGTGTGT	GTGTGTGTGT	ATATATATAT	1500
ATATATATAT	AAATATATAT	ATATAAAATA	TTCAGCAGCA	CCAAGTTTTA	TAACTATTGT	1560
TTGTTTGACT	TTATTAATAC	TAGAATATGT	AGTCTCAGCC	TTAATTTTAC	ATTTACATTA	1620
TTTTGTAATT	TTTTATTACT	ATTTTTAAGG	GGTTAAAGAG	AACATACATT	CTCACATTAG	1680
TGTACTTTCT	GGTAGAAAGT	TGCTGCAAAA	ACATTTGAAA	TGTATATTAA	CCTAATGTAT	1740
GTCATATATA	TGTCTTTGTG	TAAGTTCAAG	ACTATTGATC	TGTGAAGTTA	TTTTGTAAGG	1800
ACATACATTT	GGTAAGTAAG	TTTGTGTCCC	AGGAAATGTA	TGTGTTTTTA	AACCCTTTCT	1860
AAATATGCAG	GCCATTAATA	AATAAGATTG	TGTCTCA			1897

## (2) INFORMATION FOR SEQ ID NO: 86:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1488 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: TESTTUT02
  - (B) CLONE: 1275261
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 86:

CCCACGCGTC	CGGGGACATC	CTGTTCTGAG	TCAAGATTCC	TCCTTCTGAA	CATGGGACTT	60
TCCAGAAGGA	CCACAGCTCC	TCCCGTGCAT	CCACTCGGCC	TGGGAGGTTC	TGGATTTTGG	120
CTGTCGAGGG	AGTTTGCCTG	CCTCTCCAGA	GAAAGATGGT	CATGAGGCCC	CTGTGGAGTC	180
TGCTTCTCTG	GGAAGCCCTA	CTTCCCATTA	CAGTTACTGG	TGCCCAAGTG	CTGAGCAAAG	240
TCGGGGGCTC	GGTGCTGCTG	GTGGCAGCGC	GTCCCCCTGG	CTTCCAAGTC	CGTGAGGCTA	300
TCTGGCGATC	TCTCTGGCCT	TCAGAAGAGC	TCCTGGCCAC	GTTTTTCCGA	GGCTCCCTGG	360
AGACTCTGTA	CCATTCCCGC	TTCCTGGGCC	GAGCCCAGCT	ACACAGCAAC	CTCAGCCTGG	420
AGCTCGGGCC	GCTGGAGTCT	GGAGACAGCG	GCAACTTCTC	CGTGTTGATG	GTGGACACAA	480
GGGGCCAGCC	CTGGACCCAG	ACCCTCCAGC	TCAAGGTGTA	CGATGCAGTG	CCCAGGCCCG	540
TGGTACAAGT	GTTCATTGCT	GTAGAAAGGG	ATGCTCAGCC	CTCCAAGACC	TGCCAGGTTT	600
TCTTGTCCTG	TTGGGCCCCC	AACATCAGCG	AAATAACCTA	TAGCTGGCGA	CGGGAGACAA	660
CCATGGACTT	TGGTATGGAA	CCACACAGCC	TCTTCACAGA	CGGACAGGTG	CTGAGCATTT	720
CCCTGGGACC	AGGAGACAGA	GATGTGGCCT	ATTCCTGCAT	TGTCTCCAAC	CCTGTCAGCT	780
GGGACTTGGC	CACAGTCACG	CCCTGGGATA	GCTGTCATCA	TGAGGCAGCA	CCAGGGAAGG	840
CCTCCTACAA	AGATGTGCTG	CTGGTGGTGG	TGCCTGTCTC	GCTGCTCCTG	ATGCTGGTTA	900
CTCTCTTCTC	TGCCTGGCAC	TGGTGCCCCT	GCTCAGGGAA	AAAGAAAAAG	GATGTCCATG	960
CTGACAGAGT	GGGTCCAGAG	ACAGAGAACC	CCCTTGTGCA	GGATCTGCCA	TAAAGGACAA	1020
TATGAACTGA	TGCCTGGACT	ATCAGTAACC	CCACTGCACA	GGCACACGAT	GCTCTGGGAC	1080
ATAACTGGTG	CCTGGAAATC	ACCATGGTCC	TCATATCTCC	CATGGGAATC	CTGTCCTGCC	1140
TCGAAGGAGC	AGCCTGGGCA	GCCATCACAC	CACGAGGACA	GGAAGCACCA	GCACGTTTCA	1200
CACCTCCCCC	TTCCCTCTCC	CATCTTCTCA	TATCCTGGCT	CTTCTCTGGG	CAAGATGAGC	1260
CAAGCAGAAC	ATTCCATCCA	GGACACTGGA	AGTTCTCCAG	GATCCAGATC	CATGGGGACA	1320
TTAATAGTCC	AAGGCATTCC	CTCCCCCACC	ACTATTCATA	AAGTACTAAC	CAACTGGCAC	1380
CAAGAAAAA	TCCTCACTAA	CCGCATCATC	CGACAACTAA	TAATTCACAC	TACATCCAAA	1440
CATCACTTAG	GCGGCGGGC	CGCCGACTGG	TTCCGGGCTT	AGGGTGGG		1488

# (2) INFORMATION FOR SEQ ID NO: 87:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1357 base pairs
    (B) TYPE: nucleic acid
    (C) STRANDEDNESS: single
    (D) TOPOLOGY: linear

(vii) IMMEDIATE SOURCE:

(A) LIBRARY: COLNNOT16

(B) CLONE: 1281682

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 87:

CCGACTTTGT AGCATTTTTA TTTAAGCTAA AACAGAGCAC ATGTATATGT ACATAAGACA 60 TCTTATGACT TTGTGGTTTT ATAGATGTTC TAGAAACTTT GTATGTAGGT ATCTACAAAA 180 TTAGTTCATT CCCCTGAATA TTTTTGCATT CATATTTTTG AGGTCTTGAT GTTTTCAGCC 240 TCTGGCGAAT CTTTTTCATT GAATTTGAAC CATTTGTAAA ATCTGTGATG CTGAAGCAGA 300 GTGTGTCACA AAGTGATGAG AACATTACTA AAATCCACGG ACGCACTGCG ACCTAAGGGC 360 TCAACGGCTG ACTCGGCAGC GGGCAGCCAC CCCACGCTCC CCTGCGGTCA CTCGCACACC 420 ACAGCCTGAA GCTCCCCCAG CGCCTGCACC TCGCACACAG CTAAGGTCAA AGTTCAAACG 480 CACTCCACAC GGAAGCTCAT TCTATACCCG AAGAGCAGTC TCAGAAAGCA AGATTACTTT 540 TGTGTTTTTT AAAAAATGAT TCTTTAATGT ATTTTTCTAA ACATTCTGAT TGGAAGTAGT 600 GGATTCCTAA ATGATTCCAA AGTCATCTGT AATTCTTCTG TTTTTGTTTT GTTCTGTCTT 660 TTCTTCATTT TGGCTTTGGG TGGGGGGAGG GGCAGGTGAC ACAAAGGATT TTTTTTTTT 720 TTTTTTTTA ATTTTTGGAA TCTTTTCCAA TAACCAGCTA AAGATTTGCA CTGAAATACA 780 ACTTGTATGC CTTTTGCATT TTTAAAGCCT GCTTCCTGGA TTTAAGCAGA GTGATAGTGT 840 TCAAAGAGCC AGTTCAGCCT GTAACATATT TGAAAAAGAT ATGTCTGCAC TTTGAGGTCC 900 CTTTTGAATG CCATTCACTA GACCTCTCAA GCATTTTGTT TCATTGCTAC ATCCAAGCGC 960 CTCACAAGTC CACAATGCGG GACAGCATCA AAAGCTCAAG ACTTTGGAAA AAGCTTGTGG 1020 GCTTGCACTG GGGGAGGGAA GGGAACAAAA TTTGTGTACT TCTTTGTTTA ATTTAGAAAT 1080 AAGGCATCCA AGAGATGCCA TTATTTTCTG TGTTTCAATT GTTGTGCCTT TGAGTTAAAC 1140 TGCATTTTTG TCTTTTGGTT GAAATCTGAA ATGTACTGTC CCAATATAAA ACAGTAATTA 1200 TTTGACCTTT GCACTGTTTG TCTGGTCCTT TTCAGTTTGA TTGCATATAA ATGTGGAACT 1260 TGATAGATCT CTATATTTTT AATGCACTTG TGATAAACTG GCAGCAGGGT TAGACATTAC 1320 TTTCAAAGCT TGAGGTAGAC CGAGTCAGCA TGCTAGA 1357

### (2) INFORMATION FOR SEQ ID NO: 88:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2330 base pairs

(B) TYPE: nucleic acid

- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRSTNOT07
  (B) CLONE: 1298305
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 88:

CCTACTTGTT	CCCACCTTGG	GAGAGGACGA	TGACTTGGGA	GGGACGCGTG	AAGGGAGAAG	60
GGGTCCTCCC	ATGAGGCTGA	GGATGGCCTG	AACCTGGAGC	AGCGGACCAG	GCAGACGGGC	120
TGAAGTGGGG	TCCCAAATTC	CATGTCCAGA	GGTGTGGGGA	GCCTGCCTCC	CTAGCTCCTG	180
GCCCCTGCCA	GGGGCTTACA	TCAAAACACC	TCAGAGGGCT	GCCCTCCAGA	GGCTGCACCC	240
AGAACAGTGG	GACATGAGCA	GGGGTGTGGG	CTTGGAGGGT	GAAGAGGATG	TGGTCCTATC	300
AGATGCTGGG	CCTCCTCAGC	CATAGCCCCC	TGCTCCTACC	CCCTGACTGG	CTCTTGTGTC	360
CTCACCTCTC	ACCCTCTCCT	TCCTGGGAGG	CCCTGGGAGG	TGATCATTGA	CACCCAGCCA	420
AGCAGACAGC	TGCGGGTGCC	CAAGCCCTTG	CTGGGCCTGC	GCGTGAGGAG	TCCCACTGCT	480
TCTAAAGGAA	GTCCTGGGCA	GGAGGTGGCT	TTGGTGGTTG	GTTCCAAAGT	TGAAAATGCT	540
TGCAGTTTGA	CCTTAGAAGA	AGTGGGAAGA	AGAAGGAGCT	CTACAGGGTC	AGCTTTGTTT	600
GATTTGTCCA	GTCTAAGAAG	TCCCATTGCC	AAAGCTTTCT	GCAGGAGGGT	GAATGCCGCA	660
GCTTGGCAGC	CCCTGGGTTT	CTCTTGGAAA	TGGTCAGTTT	CCCCTCAAAG	TACCCAAAGT	720
AGCCTTGGCT	TGAGTTTTTG	TCCTTGCCTC	CTTTTTAGAG	AAGAGGGCAT	TTAGACTGCA	780
TTTTCCTGGT	TAAAGAAGGT	TAAAGCAAAT	GTTTATTGCC	TTTTCTAGTG	AACTAACTCG	840
TAGAGATGTT	CTCAGCAGGA	AGACAGTCTT	AGCACTGTCA	CTTAGCAGAT	TGCACTTAAG	900
TCCCTTGTGC	TGGCCAGATG	GCGTGGCTGG	TTGCCTTAAT	ATGTCCCAGG	ACCCCTGACA	960
GGGCTGCCTG	GCCTCTCCCT	CGTGCTCCTC	AAGAGCCCAG	TCCATACACT	GTGGATGTCA	1020
TTGCTGTCGG	GTTAGGAAGT	CTTGTCCTAG	AACGCCCTGG	CTGGTATGAC	CACAGTTCAT	1080
GGCGGCTCTT	CTCGCTTGGG	TCATGGTCAT	CTTCCAGCAC	CTGCTGTGCT	GGGAAGGCCG	1140
AGGATGGGGG	CCCAGCACTG	TCCAGGCCTG	CTGGGGCCTG	GCTGGGAGTC	CTGTGGGCAG	1200
CATGGAACAT	GCAGCTGGGC	TTCCTGTGAC	CAGGCACCCT	CTGGCACTGT	TGCTTGCCCT	1260
GTGCCCTGGA	CCTTTTCCTG	CCCTTCTCCT	TCCTCTGCTC	CCTTGGGGCT	ACCCCTTGGC	1320
CCCTCCTGGT	CTGTGCAAAC	TCCCTCAGGG	AGCCCCCCTG	CCCTGTAGCT	CTCACTTAAC	1380
TTCCTAGGGG	CTGCTGAGCC	CACCCAGAGG	TTGTTGGAGT	TCAGCGGGGC	AGCTTGTCTC	1440
CCTTGTCAGC	AGGGGCGTAA	GGGCTGGGTT	TGGCCATACA	AGGTTGGCTA	CGCCCTCAAT	1500
CCCTGACCGT	TCCAGGCACT	GAGCTGGGCA	CCCACGGAAG	GACATGCTGT	CCAGACTGTG	1560

#### (2) INFORMATION FOR SEQ ID NO: 89:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2729 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LUNGNOT12
  - (B) CLONE: 1360501
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89:

CTACACCTTT TCCATTTGCT AATAAGGCCC TGCCAGGCTG GGAGGGAATT GTCCCTGCCT 60
GCTTCTGGAG AAAGAAGATA TTGACACCAT CTACGGGCAC CATGGAACTG CTTCAAGTGA 120
CCATTCTTTT TCTTCTGCCC AGTATTTGCA GCAGTAACAG CACAGGTGTT TTAGAGGCAG 180
CTAATAATTC ACTTGTTT ACTACAACAA AACCATCTAT AACAACACCA AACACAGAAT 240
CATTACAGAA AAATGTTGTC ACACCAACAA CTGGAACAAC TCCTAAAGGA ACAATCACCA 300
ATGAATTACT TAAAATGTCT CTGATGTCAA CAGCTACTTT TTTAACAAGT AAAGATGAAG 360
GATTGAAAGC CACAACCACT GATGTCAGA AGAATGACTC CATCATTTCA AACGTAACAG 420
TAACAAGTGT TACACTTCCA AATGCTGTTT CAACATTACA AAGTTCCAAA CCCAAGACTG 480
AAACTCAGAG TTCAATTAAA ACAACAGAAA TACCAGGTAG TGTTCTACAA CCCAGATGCAT 540



CACCTTCTAA	AACTGGTACA	TTAACCTCAA	TACCAGTTAC	AATTCCAGAA	AACACCTCAC	600
AGTCTCAAGT	AATAGGCACT	GAGGGTGGAA	AAAATGCAAG	CACTTCAGCA	ACCAGCCGGT	660
CTTATTCCAG	TATTATTTTG	CCGGTGGTTA	TTGCTTTGAT	TGTAATAACA	CTTTCAGTAT	720
TTGTTCTGGT	GGGTTTGTAC	CGAATGTGCT	GGAAGGCAGA	TCCGGGCACA	CCAGAAAATG	780
GAAATGATCA	ACCTCAGTCT	GATAAAGAGA	GCGTGAAGCT	TCTTACCGTT	AAGACAATTT	840
CTCATGAGTC	TGGTGAGCAC	TCTGCACAAG	GAAAAACCAA	GAACTGACAG	CTTGAGGAAT	900
TCTCTCCACA	CCTAGGCAAT	AATTACGCTT	AATCTTCAGC	TTCTATGCAC	CAAGCGTGGA	960
AAAGGAGAAA	GTCCTGCAGA	ATCAATCCCG	ACTTCCATAC	CTGCTGCTGG	ACTGTACCAG	1020
ACGTCTGTCC	CAGTAAAGTG	ATGTCCAGCT	GACATGCAAT	AATTTGATGG	AATCAAAAAG	1080
AACCCCGGGG	CTCTCCTGTT	CTCTCACATT	TAAAAATTCC	ATTACTCCAT	TTACAGGAGC	1140
GTTCCTAGGA	AAAGGAATTT	TAGGAGGAGA	ATTTGTGAGC	AGTGAATCTG	ACAGCCCAGG	1200
AGGTGGGCTC	GCTGATAGGC	ATGACTTTCC	TTAATGTTTA	AAGTTTTCCG	GGCCAAGAAT	1260
TTTTATCCAT	GAAGACTTTC	CTACTTTTCT	CGGTGTTCTT	ATATTACCTA	CTGTTAGTAT	1320
TTATTGTTTA	CCACTATGTT	AATGCAGGGA	AAAGTTGCAC	GTGTATTATT	AAATATTAGG	1380
TAGAAATCAT	ACCATGCTAC	TTTGTACATA	TAAGTATTTT	ATTCCTGCTT	TCGTGTTACT	1440
TTTAATAAAT	AACTACTGTA	CTCAATACTC	TAAAAATACT	ATAACATGAC	TGTGAAAATG	1500
GCAATGTTAT	TGTCTTCCTA	TAATTATGAA	TATTTTTGGA	TGGATTATTA	GAATACATGA	1560
ACTCACTAAT	GAAAGGCATT	TGTAATAAGT	CAGAAAGGGA	CATAGGATTC	ACATATCAGA	1620
CTGTTAGGGG	GAGAGTAATT	TATCAGTTCT	TTGGTCTTTC	TATTTGTCAT	TCATACTATG	1680
TGATGAAGAT	GTAAGTGCAA	GGGCATTTAT	AACACTATAC	TGCATTCATT	AAGATAATAG	1740
GATCATGATT	TTTCATTAAC	TCATTTGATT	GATATTATCT	CCATGCATTT	TTTATTTCTT	1800
TTAGAAATGT	AATTATTTGT	TCTAGCAATC	ATTGCTAACC	TCTAGTTTGT	AGAAAATCAA	1860
CACTTTATAA	ATACATAATT	ATGATATTAT	TTTTCATTGT	ATCACTGTTC	TAAAAATACC	1920
ATATGATTAT	AGCTGCCACT	CCATCAGGAG	CAAATTCTTC	TGTTAAAAGC	TAACTGATCA	1980
ACCTTGACCA	CTTTTTTGAC	ATGTGAGATC	AAAGTGTCAA	GTTGGCTGAG	GTTTTTTGGA	2040
AAGCTTTAGA	ACTAATAAGC	TGCTGGTGGC	AGCTTTGTAA	CGTATGATTA	TCTAAGCTGA	2100
TTTTGATGCT	AAATTATCTT	AGTGATCTAA	GGGGCAGTTT	AGTGAAGATG	GAATCTTGTA	2160
TTTAAAATAG	CCTTTTAAAA	TTTGTTTTGT	GGTGATGTAT	TTTGACAACT	TCCATCTTTA	2220
GGAGTTATAT	AATCACCTTG	ATTTTAGTTT	CCTGATGTTT	GGACTATTTA	TAATCAAGGA	2280
CACCAAGCAA	GCATAAGCAT	ATCTATATTT	CTGACTGGTG	TCTCTTTGAG	AAGGATGGGA	2340
AGTAGAAAAA	AAAAAAAGAA	AGAAAGGAAA	GGAAGAGAGG	AGAGAAGAAG	GCAGGGATCT	2400



CCACTATGTA TGTTTCACT TTAGAACTGT TGAGCCCATG CTTAATTTA ATCTAGAAGT 2460
CTTTAAATGG TGAGACAGTG ACTGGAGCAT GCCAATCAGA GAGCATTTGT CTTCAGAAAA 2520
AAAAAAAAATC TGAGTTTGAG ACTAGCCTGG CCAACATGTT GAAACCCCAT ATCTACTAAA 2580
AATACAAAAA TTAGCCTGGT GTGGTGGCGC ACGCCTGTAG TCCCAGCTAC TCTGGAGCCT 2640
GAGGAACGTG AATCGCTTGA ACCCAGAAGA CAGGGTTGC AGTGAGCTGA GATGGCACTA 2700
TTGCACTCCA GACTGGTGAC ACACGCAGA

#### (2) INFORMATION FOR SEQ ID NO: 90:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1386 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LUNGNOT12
  - (B) CLONE: 1362406
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 90:

GGCCCCTGCA	CTGCTCCTGA	TCCCTGCTGC	CCTCGCCTCT	TTCATCCTGG	CCTTTGGCAC	60
CGGAGTGGAG	TTCGTGCGCT	TTACCTCCCT	TCGGCCACTT	CTTGGAGGGA	TCCCGGAGTC	120
TGGTGGTCCG	GATGCCCGCC	AGGGATGGCT	GGCTGCCCTG	CAGACCGCAG	CATCCTTGCC	180
CCCCTGGCAT	GGGATCTGGG	GCTCCTGCTT	CTATTTGTTG	GGCAGCACAG	CCTCATGGCA	240
GCTGAAAGAG	TGAAGGCATG	GACATCCCGG	TACTTTGGGG	TCCTTCAGAG	GTCACTGTAT	300
GTGGCCTGCA	CTGCCCTGGC	CTTGCAGCTG	GTGATGCGGT	ACTGGGAGCC	CATACCCAAA	360
GGCCCTGTGT	TGTGGGAGGC	TCGGGCTGAG	CCATGGGCCA	CCTGGGTGCC	GCTCCTCTGC	420
TTTGTGCTCC	ATGTCATCTC	CTGGCTCCTC	ATCTTTAGCA	TCCTTCTCGT	CTTTGACTAT	480
GCTGAGCTCA	TGGGCCTCAA	ACAGGTATAC	TACCATGTGC	TGGGGCTGGG	CGAGCCTCTG	540
GCCCTGAAGT	CTCCCCGGGC	TCTCAGACTC	TTCTCCCACC	TGCGCCACCC	AGTGTGTGTG	600
GAGCTGCTGA	CAGTGCTGTG	GGTGGTGCCT	ACCCTGGGCA	CGGACCGTCT	CCTCCTTGCT	660
TTCCTCCTTA	CCCTCTACCT	GGGCCTGGCT	CACGGGCTTG	ATCAGCAAGA	CCTCCGCTAC	720
CTCCGGGCCC	AGCTACAAAG	AAAACTCCAC	CTGCTCTCTC	GGCCCCAGGA	TGGGGAGGCA	780
GAGTGAGGAG	CTCACTCTGG	TTACAAGCCC	TGTTCTTCCT	CTCCCACTGA	ATTCTAAATC	840
CTTAACATCC	AGGCCCTGGC	TGCTTCATGC	CAGAGGCCCA	AATCCATGGA	CTGAAGGAGA	900
TGCCCCTTCT	ACTACTTGAG	ACTTTATTCT	CTGGGTCCAG	CTCCATACCC	TAAATTCTGA	960

GTTTCAGCCA CTGAACTCCA AGGTCCACTT CTCACCAGCA AGGAAGAGTG GGGTATGGAA 1020 GTCATCTGTC CCTTCACTGT TTAGAGCATG ACACTCTCCC CCTCAACAGC CTCCTGAGAA 1080 GGAAAGGATC TGCCCTGACC ACTCCCCTGG CACTGTTACT TGCCTCTGCG CCTCAGGGGT 1140 CCCCTTCTGC ACCGCTGGCT TCCACTCCAA GAAGGTGGAC CAGGGTCTGC AAGTTCAACG 1200 GTCATAGCTG TCCCTCCAGG CCCCAACCTT GCCTCACCAC TCCCGGCCCT AGTCTCTGCA 1260 CCTCCTTAGG CCCTGCCTCT GGGCTCAGAC CCCAACCTAG TCAAGGGGAT TCTCCTGCTC 1320 TAACTCGAT GACTTGGGGC TCCCTGCTC CCCGAGGAAG ATGCTCTGCA GGAAAATAAA 1380 AGTCAG

#### (2) INFORMATION FOR SEQ ID NO: 91:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 542 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LATRTUT02
  - (B) CLONE: 1405329
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 91:

CCCGGGCCAT	GCAGCCTCGG	ccccgcggc	GCCCGCCGCG	CACCCGAGGA	GATGAGGCTC	60
CGCAATGGCA	CCTTCCTGAC	GCTGCTGCTC	TTCTGCCTGT	GCGCCTTCCT	CTCGCTGTCC	120
TGGTACGCGG	CACTCAGCGG	CCAGAAAGGC	GACGTTGTGG	ACGTTTACCA	GCGGGAGTTC	180
CTGGCGCTGC	GCGATCGGTT	GCACGCAGCT	GAGCAGGAGA	GCCTCAAGCG	CTCCAAGGAG	240
CTCAACCTGG	TGCTGGACGA	GATCAAGAGG	GCCGTGTCAG	AAAGGCAGGC	GCTGCGAGAC	300
GGAGACGGCA	ATCGCACCTG	GGGCCGCCTA	ACAGAGGACC	CCCGATTGAC	GCCGTGGAAC	360
GGCTCACACC	GGCACGTGCT	GCACCTGCCC	ACCGTCTTCC	ATCACCTGCC	ACACCTGCTG	420
GCCAAGGAGA	GCAGTCTGCA	GCCCGCGGTG	CGCGTGGGCC	AGGGCCGCAC	CGGAGTGTCG	480
GTGGTGATGG	GCATCCCGAG	CGTGCGGCGC	GAGGTGCACT	CGTACCTGAC	TGACACTCTG	540
CA						542

#### (2) INFORMATION FOR SEQ ID NO: 92

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 772 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRAINOT12
- (B) CLONE: 1415223
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 92:

CGAGCCCGGA	GTGCGGACAC	CCCCGGGATG	CTTGCGCCCC	AGAGGACCCG	CGCCCCAAGC	60
CCCCGCGCCG	CCCCCAGGCC	CACCCGGAGC	ATGCTGCCTG	CAGCCATGAA	GGGCCTCGGC	120
CTGGCGCTGC	TGGCCGTCCT	GCTGTGCTCG	GCGCCCGCTC	ATGGCCTGTG	GTGCCAGGAC	180
TGCACCCTGA	CCACCAACTC	CAGCCATTGC	ACCCCAAAGC	AGTGCCAGCC	GTCCGACACG	240
GTGTGTGCCA	GTGTCCGAAT	CACCGATCCC	AGCAGCAGCA	GGAAGGATCA	CTCGGTGAAC	300
AAGATGTGTG	CCTCCTCCTG	TGACTTCGTT	AAGCGACACT	TTTTCTCAGA	CTATCTGATG	360
GGGTTTATTA	ACTCTGGGAT	CTTAAAGGTC	GACGTGGACT	GCTGCGAGAA	GGATTTGTGC	420
AATGGGGCGG	CAGGGGCAGG	GCACAGCCCC	TGGGCCCTGG	CCGGGGGGCT	CCTGCTCAGC	480
CTGGGGCCTG	CCCTCCTCTG	GGCTGGGCCC	TGATGTCTCC	TGCTTCCCAC	GGGGCTTCTG	540
AGCTTGCTCC	CCTGAGCCTG	TGGCTGCCCT	CTCCCCAGCC	TGGCGTGGCT	GGGGCTGGGG	600
GCAGCCTTGG	GCCAGCTCCG	TGGCTGTGGC	CTGTGGGTCT	GAATTCTTCC	CCGACGTGAA	660
GCCTNCCTGT	CTCTCCGGCA	GCTCTGAGTC	CCAGGCAGCT	GGACATTCCA	GGGGAACAAG	720
CCATTNGGCA	GGAGGGCTGG	GATGAGGTTG	GGGGGGACCG	GAGGTCCCGG	AG	772

#### (2) INFORMATION FOR SEQ ID NO: 93:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1738 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: BRAINOT12
  - (B) CLONE: 1416553
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 93:

TGTCCATCCA AAAACCATAA AATCACTGGG TTCCACATCA GCCTCCATGA GGCCAAGCCT 60
TGTACCTGCA AGCTCTTGGC CTAACCATTC CTCTGTCCTC TTCTCTGGCC TGCCTGGGGA 120
GCCCGTGAAG GCCGCACGGG TGCCTCCAGC CTGAGACATC AGGGGAGAGC CTGCAGCTGA 180

-----

#### PF-0459 US

GTTCAGCAGA	AAGGAGGAAT	CCTGGCCCTC	AGGAAGAAGA	TAGTCACATG	TTTTTCTTCC	240
TTGTCCCCAC	AGCCCCCAGA	ACAACATTCT	CCCTGCTGGC	AGCCCTTCCA	TGTCTCCAAA	300
CCTGGGTCAG	AGTGAAAGGA	CCTTTGGGGG	TGGGTGGGAG	CAAAGGGCCC	ACCTGCTGGT	360
TGGTGAAAGC	AGTGGTGCCG	GAGTGCTAGG	TACCGCACGA	GTAGTGGTGC	GGGGGCTTGG	420
GAAGCAGACC	AGGGTTGGAC	AAAACCCCAT	GAGGGCGGG	AGCTGGAAGA	AAAGTCTCTT	480
GGGGACCTCT	GGGGCAAGGA	GCTGAGAAGT	CCTGCAGCAC	CAGGTGAGAC	TTGCTTACAG	540
TGGATGCCAC	TTCTAGGCCT	CTGGACCGCA	GATGCCCTCC	TCCCTCCTGC	ACACCTGGCC	600
TCCTGGGCCT	CCAGGTAAAG	AGAGAGAGCC	AGCCCAGCCC	TGTTTCCCCT	CAGTCCTCCT	660
TTGCTCCTGC	TGCTTCTCCC	AACAGCCCAC	TGTTAGGAGG	TAGTAGACCC	CAGCCTCAAG	720
GCTCTGACCT	TCTTCATGTG	GGCACAGAGG	GTCCTGACAC	TCTGGCAGGG	CCTGAGCTGG	780
GGCAGGCCTC	CCTCAGGGCC	AGGGGCGATG	GCACCCGGG	GACAGGCAGA	CCTCCTTCCT	840
GCCGTCAGCA	CCCCCTTCCT	TATCACTGTC	TGGTCTCCGA	GCTTCGGCTG	CAGCCTGAGG	900
TGTGTCCTGG	GCTCCTCAGA	GCCTGAAGCA	AGCTTTTGGA	AGCCTGCAGT	CCTCCCAGCT	960
CCAGTGCAGA	AGCCTCTCTC	TCCAGCCTTT	CCCCAGGCAG	GAGTTGGGGT	TGGGGGCCTC	1020
TGTCCCTCAT	CGCTTACCTT	GGAAAGGTGG	GAAGCTGGCA	ATCTGCACCT	TGGGGCCTGG	1080
GCTCCCCCTC	TCTGTGCCAG	CGGCTTCCCA	GCACCTGGGA	GGGGCTGCAG	CCCCAGCTGG	1140
ACTCCAGCCT	GTCCCTCTTA	GCACTCTAGC	TGCCCACTCC	AGGGCAGGGA	CTCGAAACCC	1200
CCTCCGTCCT	GAGCAGCCAC	CTCCAGGGCC	CTGTTTGGGA	CCACTCTCTC	AGTCCCCAGG	1260
TCCTCAGGGC	CCCAGAGCGG	GAGGGTCTCC	TACCTGGAAG	TCCCCCTGAG	CTCCAGGGCC	1320
CAGCCCTACC	TGCCAGTGCT	GGTGTCAGGG	CACTCAACAC	CGAGTGTGGG	GGCCACGCCC	1380
CTTGCCATGC	CCACGGCCTC	CTCCTGTAGC	CCCTGCCTGC	ACCCACGATG	CTGCACGGGC	1440
CCGCCCTGGT	GGGGCTCGGC	GAGTAATGTG	TTTTGTCCCC	AGTTAACCAC	CATTCTGCGG	1500
CCTGGTTCTG	CAAGGAACCA	GGGCTGCCCC	ACCGCCCGCC	GTCTGCCGCC	CTAGGCTTCC	1560
TGACTCCATT	AGTTCCGACA	CTTGTGAAAC	TCCGAGAAGT	GCTGTGGTCT	CAGCAATGCA	1620
CCTGTTTTGT	ACATGATTGT	GTAATTTAAA	GGTATATAAA	TACAAATATA	TATATATATC	1680
AGTTGTGATT	GTATGACTGT	GGATAAAATC	CAGAACTGTG	TCAACCTGAA	AAAAAAA	1738

#### (2) INFORMATION FOR SEQ ID NO: 94:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2100 base pairs (B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

(A) LIBRARY: KIDNNOTO9

(B) CLONE: 1418517

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 94:

GGGAAAGCGG CGAGTAAGAT GGAAGATGAG GAGGTCGCTG AGAGCTGGGA AGAGGCGGCA 60 GACAGCGGGG AAATAGACAG ACGGTTGGAA AAAAAACTGA AGATCACACA AAAAGAGAGC 120 AGGAAATCCA AATCTCCTCC CAAAGTGCCC ATTGTGATTC AGGACGATAG CCTTCCCGCG 180 GGGCCCCCTC CACAGATCCG CATCCTCAAG AGGCCCACCA GCAACGGTGT GGTCAGCAGC 240 CCCAACTCCA CCAGCAGGCC CACCCTTCCA GTCAAGTCCC TAGCACAGCG AGAGGCCGAG 300 TACGCCGAGG CCCGGAAGCG GATCCTGGGC AGCGCCAGCC CCGAGGAGGA GCAGGAGAAA 360 CCCATCCTCG ACAGGCCAAC CAGGATCTCC CAACCCGAAG ACAGCAGGCA GCCCAATAAT 420 GTGATCAGAC AGCCTTTGGG TCCTGATGGG TCTCAAGGCT TCAAACAGCG CAGATAAATG 480 CAGGCAAGAA AAGATGCCGC CGTTGCTGCC GTCACCGCCT CCTGGGTCGT CCGCCACGGG 540 TTGCACTGCC GTGGCAGACA GCTGGACTTG AGCAGAGGGA ACGACCTGAC TTACTTGCAC 600 TGTGATCCCC CTTGCTCCGC CCACTGTGAC CTTGAACCCC ATGCACTGTG ACCTCCCCCC 660 TTCTCCCCCT TCCCACTGTG ATTGGCACAT CGACAAGGGC TGTCCCAAGT CAATGGAAAG 720 GGAAAGGGTG GGGGTTAGGG GAAGGTTGGG GGGACCCAGC AAGGACTCAG AGAGTCAGAC 780 AGTGCCACTT GGCCACTTGG GGTAAAGCCA GTGCCAGCAA TAACAGTTTA TCATGCTCAT 840 TAATTTGGGA TTTCAAAACA CAAATGAAAA CTCACACCCA CCCACCCCCA AGTGCATGTC 900 TCCATCACTT AAAAAGTAAG TTCCATTTGA AAATATCCTT TCTTTTTTT TTCTTCCTAT 960 TTTTGTTTGT TTATACAAAT ATCTGATTTG CAAGAAAAAG TGCATGGGAG GGGTTTTAGT 1020 GGTTTAATGA ATTTTTAATT AAGAAAGGGT AGTTTGGTAG TCTACTTAAA AATGTTTCTG 1080 GGAAATTCAC TAGAAACATT AACCAATAGG ATTTTGGTGA GCTTAGCTTC TGTATTCCTA 1140 CTGCCGCCCA GAAAAGGGGC AGGGCTCTGC AGCCGCCAGG ACAGACGAGC ACCCCATGCC 1200 TATACCTCCC TCCCCGAGCT AAGTCCCAGG GCATCTGGGC CTTGCCTGGA GACTGGGCTA 1260 GCTCTGTAGG CTCGGAGAGC CTGGGGAGGG TGCCAACCCC ACCTCTAGTA TTTTGGGAGA 1320 TAGGGAAAGT GAACCGACTT CCCCTTCCCA TACCCCTCAG GGTGGTTCCC TACCAGCCAG 1380 GCTTACTACT TCTAGAAGAA AGCAGAGTGC CAGGGAGTGA GATTGCATCC CTGGGCTTAG 1440 AAGTGACGGA GAGAAGACTT GTTTAGTATT TTGCCATCAG CACAAGGAAA ACCAGGAGAG 1500 AGTCTGCCTC CAGGACTCTG AGCCTTCTGC CTCGTATGTT CAGAAGGTGG ATAGGTCTTC 1560

CCACTCCAGC ATGGCTTGAA CTCTTAGGGG TCTGCAGTGC TCCATCTCCA TTGGTGGCCC 1620
CAGCTCAGTA ACTATACCTG GTACATTTCC TGTGTGCAAT CAGTACCTTG AAGGCAGAAC 1680
ATTCTGAATA AAGTTGGAAA AAGAACAGCT TTGCTTTGCA AAGATTGATG ACAGACTGGT 1740
TCCTCAGAGG CCTAGGCTAC CCGTCACCCC TTTTTCCAGA GCGAGGGCCT GGAATGAAGG 1800
CAGTTTATCC TCTGTCCCTG GAGCCTGGGG TTTGCTTTGG CTCCTTGAGG TGGAAGAGAC 1860
TAAGAGGGCA GCTGCCCAGA GCAGCTGTGT GTACCTGGCT CCTCTCAGGC TTCCTGATCC 1920
CTTCCATTGC ACTGCGCTT ATCCCTCAGC CAGCCAGACA GCCTCCCTGC TCCTGACCAG 1980
CAGATACGTT TCGGAGTGGT TGGTGTGTT TTTGTGATGA GGGCAGCACA TGGTGGCCAA 2040
GGTGGGCAAA GCTGAGTCTC ACAAGGCTCA AATCCCTTCG GTTGGGNTCC CCTTGTGGGG 2100

#### (2) INFORMATION FOR SEQ ID NO: 95:

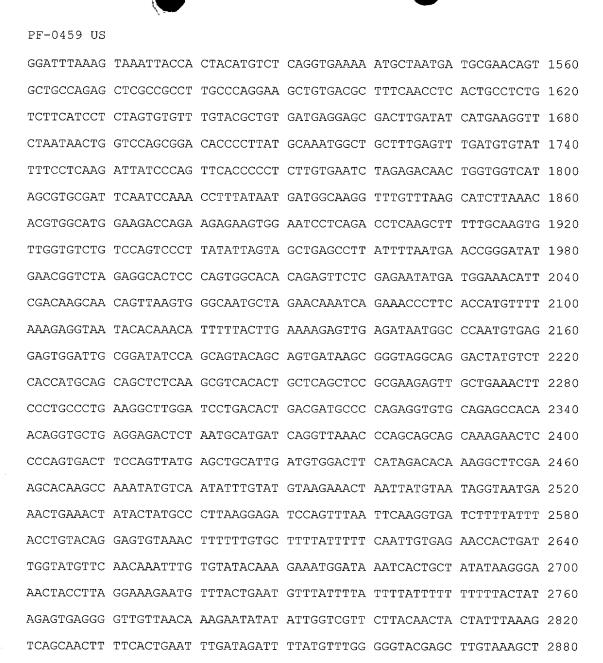
- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2458 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: PANCNOTO8
  - (B) CLONE: 1438165
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 95:
- GCGGGCGGAG ATGTAGACCC GGTAGTGTTG TGCCTTGTGG TGACAACTGG CGGCAGCGCG 60
  CCGCGGGCCC GAGACTTAGT CTCGGGCCGC CATGGCCAGC GTCCACGAGA GCCTCTACTT 120
  CAATCCCATG ATGACCAATG GGGTTGTGCA CGCCAATGTG TTCGGCATCA AGGACTGGGT 180
  GACGCCGTAC AAGATCGCGG TGCTGGTGCT GCTGAACGAG ATGAGCCGCA CAGGCGAGGG 240
  CGCCGTCAGC CTCATGGAGC GGCGGAGGCT CAACCAGCTG CTCCTGCCCC TGCTGCAGGG 300
  CCCAGATATT ACACTGTCAA AACTTTACAA GTTAATTGAA GAGTCTTGTC CACAGCTGGC 360
  AAATTCAGTG CAGATCAGAA TCAAACTGAT GGCTGAAGGC GAGTTGAAGG ATATGGAACA 420
  GTTTTTTGAT GACCTTTCAG ATTCTTTCTC TGGAACTGAA CCAGAGGTTC ACAAAACAAG 480
  TGTAGTAGGT TTGTTTCTGC GTCACATGAT CTTGGCCTAC AGTAAGCTTT CTTTCAGCCA 540
  AGTGTTTAAA CTGTACACTG CCCTTCAGCA GTACTTCCAG AATGGTAGA AAAAGACAGT 600
  GGAGGATGCT GATATGGAAC TGACCAGTAG AGATGAGGGT GAAAGAAAAA TGGAAAAAGA 660
  AGAACTTGAT GTATCTGTAA GAGAAGAGGA GGTATCTTGC AGTGGGCCTC TGTCCCAAAA 720
  ACAAAGCAGAA TTTTTTCTTT CTCAACAGGC TTCTTTGCTA AAGAATGATG AGACTAAGGC 780

CCTCACTCCA	GCTTCCTTGC	AGAAGGAATT	AAACAATTTG	TTGAAATTTA	ATCCTGATTT	840
TGCTGAAGCG	CATTATCTCA	GCTACTTAAA	CAACCTCCGT	GTCCAAGATG	TTTTCAGTTC	900
AACACACAGT	CTCCTCCATT	ATTTTGATCG	TCTGATTCTT	ACCGGAGCCG	AAAGCAAAAG	960
TAATGGGGAA	GAGGGCTATG	GCCGGAGCTT	GAGATACGCC	GCTCTGAATC	TTGCCGCCCT	1020
GCACTGCCGC	TTCGGTCACT	ATCAACAGGC	AGAGCTCGCC	CTGCAGGAGG	CAATTAGGAT	1080
TGCCCAGGAG	TCCAACGATC	ACGTGTGTCT	CCAGCACTGT	TTGAGCTGGC	TTTATGTGCT	1140
GGGGCAGAAG	AGATCCGATA	GCTATGTTCT	GCTGGAGCAT	TCTGTGAAGA	AGGCAGTACA	1200
TTTTGGGTTA	CCGAGAGCTT	TTGCTGGGAA	GACGGCAAAC	AAGCTGATGG	ATGCCCTAAA	1260
GGACTCCGAC	CTCCTGCACT	GGAAACACAG	CCTGTCAGAG	CTCATCGATA	TCAGCATCGC	1320
ACAGAAAACG	GCCATCTGGA	GGCTGTATGG	CCGCAGCACC	ATGGCACTGC	AACAGGCCCA	1380
GATGTTGCTG	AGCATGAACA	GCCTGGAGGC	GGTGAATGCG	GGCGTGCAGC	AGAACAACAC	1440
AGAGTCCTTT	GCTGTCGCAC	TCTGCCACCT	CGCAGAGCTA	CACGCGGAGC	AGGGCTGTTT	1500
TGCTGCAGCT	TCTGAAGTGT	TAAAGCACTT	GAAGGAACGA	TTTCCGCCTA	ATAGTCAGCA	1560
CGCCCAGTTA	TGGATGCTAT	GTGATCAAAA	AATACAGTTT	GACAGAGCAA	TGAATGATGG	1620
CAAATATCAT	TTGGCTGATT	CACTTGTTAC	AGGAATCACA	GCTCTCAATA	GCATAGAGGG	1680
TGTTTATAGG	AAAGCGGTTG	TATTACAAGC	TCAGAACCAA	ATGTCAGAGG	CACATAAGCT	1740
TTTACAAAAA	TTGTTGGTTC	ATTGTCAGAA	ACTGAAGAAC	ACAGAAATGG	TGATCAGTGT	1800
CCTACTGTCC	GTGGCAGAGC	TGTACTGGCG	ATCTTCCTCC	CCTACCATCG	CGCTGCCCAT	1860
GCTCCTGCAG	GCTCTGGCCC	TCTCCAAGGA	GTACCGGTTA	CAGTACTTGG	CCTCTGAAAC	1920
AGTGCTGAAC	TTGGCTTTTG	CGCAGCTCAT	TCTTGGAATC	CCAGAACAGG	CCTTAAGTCT	1980
TCTCCACATG	GCCATCGAGC	CCATCTTGGC	TGACGGGGCT	ATCCTGGACA	AAGGTCGTGC	2040
CATGTTCTTA	GTGGCCAAGT	GCCAGGTGGC	TTCAGCAGCT	TCCTACGATC	AGCCGAAGAA	2100
AGCAGAAGCT	CTGGAGGCTG	CCATCGAGAA	CCTCAATGAA	GCCAAGAACT	ATTTTGCAAA	2160
GGTTGACTGC	AAAGAGCGCA	TCAGGGACGT	CGTTTACTTC	CAGGCCAGAC	TCTACCATAC	2220
CCTGGGGAAG	ACCCAGGAGA	GGAACCGGTG	TGCGATGCTC	TTCCGGCAGC	TGCATCAGGA	2280
GCTGCCCTCT	CATGGGGTAC	CCTTGATAAA	CCATCTCTAG	AGAGGACATC	CCTGCTGGGC	2340
TGCTGTGCAG	AGTATAAGAT	TTTGGACTTG	TTCATGTCCC	CTCTCTCCCT	ATAAATGATG	2400
TATTTGTGAC	ACCCTATCTT	GTCAATAAAC	AGCATTCTGA	TTAAAAAAAA	AAAAAAA	2458

(2) INFORMATION FOR SEQ ID NO: 96:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2900 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: THYRNOTO3
    (B) CLONE: 1440381
- (2) 0201.21 2110002
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 96:

TGCATGGATG GGATA	ACTGGA TGAATCTTTG	CTTGAAACCT	GTCCAATTCA	GTCACCATTA	60
CAAGTTTTTG CAGGA	AATGGG TGGACTGGCT	CTTATTGCTG	AAAGACTACC	CATGCTATAT	120
CCAGAAGTAA TTCAA	ACAGGT GAGTGCTCCA	GTTGTAACAT	CTACCACTCA	GGAAAAGCCG	180
TATGATAGCG ATCAC	GTTTGA ATGGGTGACO	ATTGAACAGT	CAGGGGAGTT	AGTTTATGAA	240
GCACCAGAAA CTGTT	TGCGGC TGAACCTCC	CCTATCAAGT	CAGCAGTACA	GACCATGTCT	300
CCCATACCTG CCCAT	TTCTTT GGCTGCTTTT	GGATTATTTC	TTCGTCTTCC	GGGCTATGCG	360
GAAGTGCTAC TGAAA	AGAGAG AAAACATGCO	CAGTGCCTTC	TTCGATTGGT	ATTGGGAGTG	420
ACAGATGATG GAGA	AGGAAG TCATATTCTT	CAATCTCCAT	CAGCCAATGT	GCTTCCAACC	480
CTTCCTTTCC ACGT	CCTTCG TAGCTTGTTT	AGCACTACAC	CTTTGACAAC	TGATGATGGT	540
GTACTTCTAA GGCGC	GATGGC ATTGGAAATT	GGAGCCTTAC	ACCTCATTCT	TGTCTGTCTC	600
TCTGCTTTGA GCCAC	CCATTC CCCACGAGTT	CCAAACTCTA	GCGTGAATCA	AACTGAGCCA	660
CAGGTGTCAA GCTC	ICATAA CCCTACATCA	ACAGAAGAAC	AACAGTTATA	TTGGGCCAAA	720
GGGACTGGCT TTGG	AACAGG CTCTACAGCT	TCTGGGTGGG	ATGTGGAACA	AGCCTTAACT	780
AAGCAAAGGC TGGAA	AGAGGA ACATGTTACO	TGCCTTCTGC	AGGTTCTTGC	CAGTTACATA	840
AATCCCGTCA GTAG	IGCGGT AAATGGAGAA	GCTCAGTCAT	CTCATGAGAC	TAGAGGGCAG	900
AACAGTAATG CCCTT	CCTTC TGTACTTCTC	C GAGCTTCTCA	GTCAGTCCTG	CCTCATCCCA	960
GCCATGTCAT CTTAT	ICTACG AAATGATTCA	GTTCTGGACA	TGGCAAGACA	TGTGCCACTC	1020
TATCGGGCAC TGCT	GGAATT GCTTCGGGCC	ATTGCTTCTT	GTGCTGCCAT	GGTGCCCCTA	1080
TTGTTGCCCC TTTC	FACAGA GAACGGTGAA	GAGGAAGAAG	AACAGTCAGA	ATGTCAAACT	1140
TCTGTTGGTA CATTO	GTTAGC CAAAATGAAG	ACCTGTGTTG	ATACCTATAC	CAACCGTTTA	1200
AGATCTAAAA GGGAA	AAATGT TAAAACAGGA	GTAAAACCAG	ATGCGTCTGA	TCAAGAACCA	1260
GAAGGACTTA CTCT	ITTGGT ACCAGACATO	CAAAAGACTG	CTGAGATAGT	TTATGCAGCC	1320
ACCACCAGTT TGCG	GCAAGC AAATCAGGAA	AAAAACTGGG	TGAATACTCC	AAGAAGGCGG	1380
CTAATGAACC CCAA	ACCTTT GTCAGTATTA	AAGTCACTTG	AAGAAAAATA	TGTGGCTGTT	1440
ATGAAGAAAT TACAC	GTTTGA TACGTTTGAA	ATGGTTTCTG	AAGATGAAGA	TGGGAAATTG	1500



#### (2) INFORMATION FOR SEQ ID NO: 97

CGGGTGCCTN ATGAGTGACC

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1310 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

2900

(A) LIBRARY: LUNGNOT14

(B) CLONE: 1510839

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 97 :

CCGCTGAGAT	GTACGAACTT	CCGGTTCTCC	GGGCAGCTGC	CACTGCTGTA	GCTTCTGCCA	60
CCTGCCACGA	CCGGGCCTCT	CCCTGGCGTT	TGGTCACCTC	TGCTTCATTC	TCCACCGCGC	120
CTATGGTCCC	TCTTGGAGCC	AGCGTGGCGG	GCCTGGCGGC	TCCCGGGTGG	TGAGAGAGCG	180
GTCCGGGAAC	GATGAAGGCC	TCGCAGTGCT	GCTGCTGTCT	CAGCCACCTC	TTGGCTTCCG	240
TCCTCCTCCT	GCTGTTGCTG	CCTGAACTAA	GCGGGCCCCT	GGCAGTCCTG	CTGCAGGCAG	300
CCGAGGCCGC	GCCAGGTCTT	GGGCCTCCTG	ACCCTAGACC	ACGGACATTA	CCGCCGCTGC	360
CACCGGGCCC	TACCCCTGCC	CAGCAGCCGG	GCCGTGGTCT	GGCTGAAGCT	GCGGGGCCGC	420
GGGGCTCCGA	GGGAGGCAAT	GGCAGCAACC	CTGTGGCCGG	GCTTGAGACG	GACGATCACG	480
GAGGGAAGGC	CGGGGAAGGC	TCGGTGGGTG	GCGGCCTTGC	TGTGAGCCCC	AACCCTGGCG	540
ACAAGCCCAT	GACCCAGCGG	GCCCTGACCG	TGTTGATGGT	GGTGAGCGGC	GCGGTGCTGG	600
TGTACTTCGT	GGTCAGGACG	GTCAGGATGA	GAAGAAGAAA	CCGAAAGACT	AGGAGATATG	660
GAGTTTTGGA	CACTAACATA	GAAAATATGG	AATTGACACC	TTTAGAACAG	GATGATGAGG	720
ATGATGACAA	CACGTTGTTT	GATGCCAATC	ATCCTCGAAG	AAGAGAATGT	GCCTTTTGAT	780
GAAAGAACTT	TATCTTTCTA	CAATGAAGAG	TGGAATTTCT	ATGTTTAAGG	AATAAGAAGC	840
CACTATATCA	ATGTTGGGGG	GGTATTTAAG	TTACATATAT	TTTAACAACC	TTTAATTTGC	900
TGTTGCAATA	AATACCGTAT	CCTTTTATTA	TATCTTTATA	TGTATAGAAG	TACTCTATTA	960
ATGGGCTCAG	AGATGTTGGG	GATAAAGTAT	ACTGTAATAA	TTTATCTGTT	TGAAAATTAC	1020
TATAAAACGG	TGTTTTCTGA	TCGGTTTTTG	TTTCCTGCTT	ACCATATGAT	TGTAAATTGT	1080
TTTATGTATT	AATCAGTTAA	TGCTAATTAT	TTTTGCTGAT	GTCATATGTT	AAAGAGCTAT	1140
AAATTCCAAC	AACCAACTGG	TGTGTAAAAA	TAATTTAAAA	TTTCCTTTAC	TGAAAGGTAT	1200
TTCCCATTTT	TGTGGGGAAA	AGAAGCCAAA	TTTATTACTT	TGTGTTGGGG	TTTTTAAAAT	1260
ATTAAGAAAT	GTCTAAGTTA	TTGTTTGCAA	AACAATAAAT	ATGATTTTAG		1310

# (2) INFORMATION FOR SEQ ID NO: 98:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2272 base pairs

  - (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear

(vii) IMMEDIATE SOURCE:

(A) LIBRARY: SPLNNOTO4

(B) CLONE: 1534876

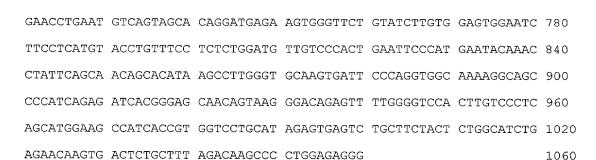
(xi) SEQUENCE DESCRIPTION: SEO ID NO: 98:

CCATGCTCCA GGCATACAGA TGTGGTTTCT CGGCTGCACC GGGCCAGGCT GCGGGTGTGC AGGCGTCTGC AAAGTTGTGC CATGTATCAG CACAGGCTTT GAGACGTCTG GACCCTGTCC 120 TTCCTCCCGT GAGGGGTTCT TGTTCTTTCT GACTCAGGTG ACTTTTCAGC CCTTCCAATT 180 CCCCTCTTTT TCTGCCCTCC CCTCCAACTC AGCCAACCCA GGTGTGGGCA GTCAGGGAGG 240 GAGGGAGTGT CCCACCACGT TCTCAGGGCA GCCCTTGACT CCTAAGCCCC TTCCTCCTTC 300 CATTCTGCAT CCCCTCCCA TCCAACCTAA ATGCCCACAG CTGGGGCTGA GCTGTATTCC 360 TGTGGAGGGA CCTCTGCCGT GCCTCTCTGA GGTCAGGCTG TGCTGTGTGA TGGGCAGGCT 420 TTGCCCCAGC CCACCCTGG CAAGGTGCAC TTGTTTTCTG GTTTGTACAA GGTGTCCTGG 480 GGGCCCGTCG CTTCCCTGCC AGTGAGGAGT GACTTCTCCC TCTCTTCCAG TCCTGTAGGG 540 GAGACAAAAC CAGATTGGGG GGCCCAAGGG GAGCATGGAA AAGGCCGGCT CCCCTGTCTT 600 TCCTTGGCTG TCAGAGTCAG GGTAACACAC ACCAAGAGTG GAGTGCGGCC AGCAAGTTTG 660 AGACCTGCCC GCCCTCCTCG CAGCTCTGCT CTGTGTCCTC AGGAAGTCAC AGAGTCTACT 720 GAGGCAAGGA GAGGGTGATT CTTTCCCCAA ATCCCTTCTT CCCTGGTTCC CAAACCAAAG 780 ACAGCCTGCA GCCCTTTCTG CATGGGGTGC TCTGTTGACA GGCTTCCCAG ATCCCTGAGT 840 CTCTCTTTCC TTCCTCCTC ATCTTTAGTT GTCCACGGTC AATTCAGTGC TTCCATTGGG 900 GGACAGTCCC CTCCGGGATG ACCTGATTCA CCTCCAGCCC AGGGAATGGA ATCTAGAGGA 960 ATACGTGGGG TGGGTCTGGA CAAGGAGCGG CAGGAATCAC CACCCATCTC CAGCTGTGGA 1020 GCCCTGTGGA GGGGAAGGGG AAGCTTGGGG TTCAGAGGGA ACTCTTCCAG GAGAGGGGTG 1080 CCCAGCGGAG GTAAAGATGA TAGAGGGTTG TGGGGGGGTCT CTAGTTGAAT GTTTTGGCCC 1140 ATGACTTTGG AACATGGCTG GCAGCTTCCA GCAGAAGTCA CGCTCCCCAT CCCCCAGGGG 1200 ACATAGGACC TTTTTCCTGC TTCCTGGTCA CTTTCAAAGA ACTATTTGCG CAATCTGTGG 1260 GTCTGTGGAT TCACGGGGCT TTCTGTGTGG GTGCTGCAGT TGCTTTTGTC TGCAGCAGCA 1320 GGACACATCT TTCCTCTTAC TCAGCCCTTT ATGGCCCATG GGGAACTCCG TGGCTCAGGG 1380 AGAGCTGAAC TCCAGGGGTG TGACCTGGGA CAGGTGGGCC TGAGGTGCCC AGCTCAGGGC 1440 AGCCAGGTGG CTCATGGGCT GTAGTGAGCC AGCTCCCTGG GGGAAAAGGC TGTGGGCCGT 1500 TAGGACCATC CTCCAGGACA GGTGACCTCT ATGAGGTCAC CTACGGCTGT GGCCGTGCAG 1560 GCCTCCTTCC AGCCCAGAGT GGCCCAGTAG AGCAAGGCAG ACAGTGACCT CCACCCCGC 1620 AGCCCTCTTA AAAGGCCAGT ACTCTTGGGG GTGGGGGGAG GGTTTAGAAA GCATTTGCCC 1680



#### (2) INFORMATION FOR SEQ ID NO: 99:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1060 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: SPLNNOT04
  - (B) CLONE: 1559131
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 99:
- GECAACTTAG CGAGCGCAAC AGGCTGCCGC TGAGGAGCTG GAGCTGGTGG GGACTGGGCC 60
  GCAATGGACA AGCTGAAGAA GGTGCTGAGC GGGCAGGACA CGGAGGACCG GAGCGGCCTG 120
  TCCGAGGTTG TTGAGGCATC TTCATTAAGC TGGAGTACCA GGATAAAAAGG CTTCATTGCG 180
  TGTTTTGCTA TAGGAATTCT CTGCTCACTG CTGGGTACTG TTCTGCTGTG GGTGCCCAGG 240
  AAAGGGACTAC ACCTCTTCGC AGTGTTTTAT ACCTTTGGTA ATATCGCATC AATTGGGAGT 300
  ACCATCTTCC TCATGGGACC AGTGAAACAG CTGAAGCGAA TGTTTGAGCC TACTCGTTTG 360
  ATTGCAACTA TCATGGTGCT GTTGTGTTT GCACTTACCC TGTGTTCTGC CTTTTGGTGG 420
  CATAACAAGG GACTTGCACT TATCTTCTGC ATTTTGCAGT CTTTTGGCATT GACGTGGTAC 480
  AGCCTTTCCT TCATACCATT TGCAAGGGAT GCTGTGAAGA AGTGTTTTGC CGTGTGTCTT 540
  GCATAATTCA TGGCCAGTTT TATGAAGCTT TGGAAGGCAC TATGGACAGA AGCTGGTGGA 600
  CAGTTTTGTA ACTATCTCG AAACCTCTGT CTTACAGACA TGTGCCTTTT ATCTTGCAGC 660
  AATGTGTTGC TTGTGATTCG AACATTTGAG GGTTACTTTT GGAAGCAACA ATACATTCTC 720



#### (2) INFORMATION FOR SEQ ID NO: 100:

PF-0459 US

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 543 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: BLADNOT03
  - (B) CLONE: 1601473
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 100:

GCTCACAGTA GCCCGGCGC CAGGGCAATC CGACCACATT TCACTCTCAC CGCTGTAGGA 60
ATCCAGATGC AGGCCAAGTA CAGCAGCACA AGGGACATGC TGGATGATGA TGGGGACACC 120
ACCATGAGCC TGCATTCTCA AGCCTCTGCC ACAACTCGGC ATCCAGAGCC CCGGCGCACA 180
GAGCACAGGG CTCCCTCTC AACGTGGCGA CCAGTGGCCC TGACCCTGCT GACTTTGTGC 240
TTGGTGCTGC TGATAGGGCT GGCAGCCCTG GGGCTTTTGT GTAAGTCTGC GCTCTGACCT 300
GGGGGGAGGAT CCTGGTTCCA AGTTTTCAG TACTACCAGC TCTCCAATAC TGGTCAAGAC 360
ACCATTTCTC AAATGGAAGA AAGATTAGGA AATACGTCCC AAGAGTTGCA ATCTCTCAA 420
GTCCAGAATA TAAAGCTTGC AGGAAGTCTG CAGCATGTGG CTGAAAAACT CTGTCGTGAG 480
CTGTATAACA AAGCTGGAGC ACACAGGTGC AGCCCTTGTA CAGAACAATG GAAATGGCAT 540
GGA

#### (2) INFORMATION FOR SEQ ID NO: 101:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2281 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

(vii) IMMEDIATE SOURCE:

(A) LIBRARY: BRAITUT12

(B) CLONE: 1615809

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 101:

AGCTGGCTCA CCTTCCAGAT TCACCTGCAG GAGCTGCTGC AGTACAAGAG GCAGAATCCA GCTCAGTTCT GCGTTCGAGT CTGCTCTGGC TGTGCTGTGT TGGCTGTTT GGGACACTAT 120 GTTCCAGGGA TTATGATTTC CTACATTGTC TTGTTGAGTA TCCTGCTGTG GCCCCTGGTG 180 GTTTATCATG AGCTGATCCA GAGGATGTAC ACTCGCCTGG AGCCCCTGCT CATGCAGCTG 240 GACTACAGCA TGAAGGCAGA AGCCAATGCC CTGCATCACA AACACGACAA GAGGAAGCGT 300 CAGGGGAAGA ATGCACCCCC AGGAGGTGAT GAGCCACTGG CAGAGACAGA GAGTGAAAGC 360 GAGGCAGAGC TGGCTGGCTT CTCCCCAGTG GTGGATGTGA AGAAAACAGC ATTGGCCTTG 420 GCCATTACAG ACTCAGAGCT GTCAGATGAG GAGGCTTCTA TCTTGGAGAG TGGTGGCTTC 480 TCCGTATCCC GGGCCACAC TCCGCAGCTG ACTGATGTCT CCGAGGATTT GGACCAGCAG 540 AGCCTGCCAA GTGAACCAGA GGAGACCCTA AGCCGGGACC TAGGGGAGGG AGAGGAGGGA 600 GAGCTGGCCC CTCCCGAAGA CCTACTAGGC CGTCCTCAAG CTCTGTCAAG GCAAGCCCTG 660 GACTCGGAGG AAGAGGAAGA GGATGTGGCA GCTAAGGAAA CCTTGTTGCG GCTCTCATCC 720 CCCCTCCACT TTGTGAACAC GCACTTCAAT GGGGCAGGGT CCCCCCAAGA TGGAGTGAAA 780 TGCTCCCTG GAGGACCAGT GGAGACACTG AGCCCCGAGA CAGTGAGTGG TGGCCTCACT 840 GCTCTGCCCG GCACCCTGTC ACCTCCACTT TGCCTTGTTG GAAGTGACCC AGCCCCCTCC 900 CCTTCCATTC TCCCACCTGT TCCCCAGGAC TCACCCCAGC CCCTGCCTGC CCCTGAGGAA 960 GAAGAGGCAC TCACCACTGA GGACTTTGAG TTGCTGGATC AGGGGGAGCT GGAGCAGCTG 1020 AATGCAGAGC TGGGCTTGGA GCCAGAGACA CCGCCAAAAC CCCCTGATGC TCCACCCCTG 1080 GGGCCCGACA TCCATTCTCT GGTACAGTCA GACCAAGAAG CTCAGGCCGT GGCAGAGCCA 1140 TGAGCCAGCC GTTGAGGAAG GAGCTGCAGG CACAGTAGGG CTTCTTGGCT AGGAGTGTTG 1200 CTGTTTCCTC CTTTGCCTAC CACTCTGGGG TGGGGCAGTG TGTGGGGAAG CTGGCTGTCG 1260 GATGGTAGCT ATTCCACCCT CTGCCTGCCT GCCTGCCTGC TGTCCTGGGC ATGGTGCAGT 1320 ACCTGTGCCT AGGATTGGTT TTAAATTTGT AAATAATTTT CCATTTGGGT TAGTGGATGT 1380 GAACAGGGCT AGGGAAGTCC TTCCCACAGC CTGCGCTTGC CTCCCTGCCT CATCTCTATT 1440 CTCATTCCAC TATGCCCCAA GCCCTGGTGG TCTGGCCCTT TCTTTTTCCT CCTATCCTCA 1500 GGGACCTGTG CTGCTCTGCC CTCATGTCCC ACTTGGTTGT TTAGTTGAGG CACTTTATAA 1560 TTTTTCTCTT GTCTTGTGTT CCTTTCTGCT TTATTTCCCT GCTGTGTCCT GTCCTTAGCA 1620 GCTCAACCCC ATCCTTTGCC AGCTCCTCCT ATCCCGTGGG CACTGGCCAA GCTTTAGGGA 1680

GGCTCCTGGT CTGGGAAGTA AAGAGTAAAC CTGGGGCAGT GGGTCAGGCC AGTAGTTACA 1740
CTCTTAGGTC ACTGTAGTCT GTGTAACCTT CACTGCATCC TTGCCCCATT CAGCCCGGCC 1800
TTTCATGATG CAGGAGAGCA GGGATCCCGC AGTACATGCC GCCAGCACTG GAGTTGGTGA 1860
GCATGTGCTC TCTCTTGAGA TTAGGAGCTT CCTTACTGCT CCTCTGGGTG ATCCAAGTGT 1920
AGTGGGACCC CCTACTAGGG TCAGGAAGTG GACACTAACA TCTGTGCAGG TGTTGACTTG 1980
AAAAATAAAAG TGTTGATTGG CTAGAACTGC TGCCTCCCTG ACTGTGAGCT GCCTTCCACA 2040
CCCTGCACTG CACTGTGTTC TCTCCTCACC CTTAACCTGC TCACTCCAG TCTGTTCTGG 2100
CTGTTTATTA CCTTGTCA AAACAGGGCC GAAGCAAGGA TTACCTTGAC AACCCTAGCT 2160
TCTCCCTTAGC CATCTTCCTT GACAGTGTGA TCTGTTTAGT GAGATTTAGC ATGTGGAAT 2220
AAAGTATATG CAGGAGGAAA TTGCTTTGTC TTCCCAATCG GTAGAAATTC GAGACCTAGC 2280

#### (2) INFORMATION FOR SEQ ID NO: 102:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 992 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: COLNNOT19
  - (B) CLONE: 1634813
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 102:

GACAGCTTGG	CCTACAGCCC	GGCGGGCATC	AGCTCCCTTG	ACCCAGTGGA	TATCGGTGGC	60
CCCGTTATTC	GTCCAGGTGC	CCAGGGAGGA	GGACCCGCCT	GCAGCATGAA	CCTGTGGCTC	120
CTGGCCTGCC	TGGTGGCCGG	CTTCCTGGGA	GCCTGGGCCC	CCGCTGTCCA	CGCCCAAGGT	180
GTCTTTGAGG	ACTGCTGCCT	GGCCTACCAC	TACCCCATTG	GGTGGGCTGT	GCTCCGGCGC	240
GCCTGGACTT	ACCGGATCCA	GGAGGTGAGC	GGGAGCTGCA	ATCTGCCTGC	TGCGATATTC	300
TACCTCCCCA	AGAGACACAG	GAAGGTGTGT	GGGAACCCCA	AAAGCAGGGA	GGTGCAGAGA	360
GCCATGAAGC	TCCTGGATGC	TCGAAATAAG	GTTTTTGCAA	AGCTCCGCCA	CAACACGCAG	420
ACCTTCCAAG	CAGGCCCTCA	TGCTGTAAAG	AAGTTGAGTT	CTGGAAACTC	CAAGTTATCA	480
TCATCCAAGT	TTAGCAATCC	CATCAGCAGC	AGCAAGAGGA	ATGTCTCCCT	CCTGATATCA	540
GCTAATTCAG	GACTGTGAGC	CGGCTCATTT	CTGGGCTCCA	TCGGCACAGG	AGGGGCCGGA	600
TCTTTCTCCG	ATAAAACCGT	CGCCCTACAG	ACCCAGCTGT	CCCCACGCCT	CTGTCTTTTG	660

GGTCAAGTCT TAATCCCTGC ACCTGAGTTG GTCCTCCTC TGCACCCCA CCACCTCCTG 720

CCCGTCTGGC AACTGGAAAG AGGGAGTTGG CCTGATTTTA AGCCTTTTGC CGCTCCGGGG 780

ACCAGCAGCA ATCCTGGGCA GCCAGTGGCT CTTGTAGAGA AGACTTAGGA TACCTCTCTC 840

ACTTTCTGTT TCTTGCCGTC CACCCCGGGC CATGCCAGTG TGTCCCTCTG GGTCCCTCCA 900

AAACTCTGGG GTTGATGGAG ATGCCCTCC CAGGCTATGC TTTTCTATAA CTTTTAAATA 960

AACCTTGGGG GTTGATGGAG TCAAAAAAAA AA 992

#### (2) INFORMATION FOR SEQ ID NO: 103:

PF-0459 US

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1554 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: UTRSNOT06
  - (B) CLONE: 1638407
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 103:

TCGCCCAGGA	GTCATCGGAC	GCCAGAATCT	GTGTCTCCAG	AACGCTATAG	CTATGGCACC	60
TCCAGCTCTT	CAAAGAGGAC	AGAGGGTAGC	TGCCGTCGCC	GTCGGCAGTC	AAGCAGTTCT	120
GCAAATTCTC	AGCAGGGTCA	GTGGGAGACA	GGCTCCCCCC	CAACCAAGCG	GCAGCGGCGG	180
AGTCGGGGCC	GGCCCAGTGG	TGGTGCCAGA	CGGCGGCGGA	GAGGGCCCC	AGCCGCACCC	240
CAGCAGCAGT	CAGAGCCCGC	CAGACCTTCC	TCTGAAGGCA	GGTGACACTG	TGATGGGGAA	300
ACAGGCTCAG	AGAGACATCC	GGCTCCGGGT	TCGAGCAGAG	TACTGCGAGC	ATGGGCCAGC	360
CTTGGAGCAG	GGCGTGGCAT	CCCGGCGGCC	CCAGGCGCTG	GCGCGGCAGC	TGGACGTGTT	420
TGGGCAGGCC	ACCGCAGTGC	TGCGCTCAAG	GGACCTGGGC	TCTGTGGTTT	GTGACATCAA	480
GTTCTCAGAG	CTCTCCTATC	TGGACGCCTT	CTGGGGCGAC	TACCTGAGTG	GCGCCCTGCT	540
GCAGGCCCTG	CGGGGCGTGT	TCCTGACTGA	GGCCCTGCGA	GAGGCTGTGG	GCCGGGAGGC	600
TGTTCGCCTG	CTGGTCAGTG	TGGATGAGGC	TGACTATGAG	GCTGGCCGGC	GCCGCCTGTT	660
GCTGATGGCG	GAGGAAGGGG	GGCGGCGCCC	GACAGAGGCC	TCCTGATCCA	GGACTGGCAG	720
GATTGATCCC	ACCTCCAAGT	CTCCGGGCCA	CCTTCTCCTG	GGAGGACGAC	CATCTCTACC	780
CCTAGAGGAC	TGTCACTCTA	GCATCTTTGA	GGACTGCGAC	AGGACCGGGA	CAGCAGGCCC	840
CTTGACAGCC	CCTCCCACAG	GATGTGGGCT	CTGAGGCCTA	AACCATTTCC	AGCTGAGTTT	900
CCTTCCCAGA	CTCCTCCTAC	CCCCAGGTGT	GCCCCTTAG	CCTCCGGAGG	CGGGGGCTGG	960

#### (2) INFORMATION FOR SEQ ID NO: 104:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1802 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: PROSTUT08
  - (B) CLONE: 1653112
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 104:

GTCGCCGGGC	TTGCGATGAA	CTTCCGGCTG	TCAAGCTCCC	GGCCGGGCTG	ACTCAAGCGG	60
AGGCGCGCGG	AACAGTCGCC	GAGGCGATTC	CCGCCCAGGC	TCCTGTAACC	GCCAGGCAGC	120
GGCCCCGCCA	TGTCCCAGCC	CCGGACCCCA	GAGCAGGCAC	TGGATACACC	GGGGGACTGC	180
CCCCCAGGCA	GGAGAGACGA	GGACGCTGGG	GAGGGGATCC	AGTGCTCCCA	ACGCATGCTC	240
AGCTTCAGTG	ACGCCCTGCT	GTCCATCATC	GCCACCGTCA	TGATCCTGCC	TGTGACCCAC	300
ACGGAGATCT	CCCCAGAACA	GCAGTTCGAC	AGAAGTGTAC	AGAGGCTTCT	GGCAACACGG	360
ATTGCCGTCT	ACCTGATGAC	CTTTCTCATC	GTGACAGTGG	CCTGGGCAGC	ACACACAAGG	420
TTGTTCCAAG	TTGTTGGGAA	AACAGACGAC	ACACTTGCCC	TGCTCAACCT	GGCCTGCATG	480
ATGACCATCA	CCTTCCTGCC	TTACACGTTT	TCGTTAATGG	TGACCTTCCC	TGATGTGCCT	540
CTGGGCATCT	TCTTGTTCTG	TGTGTGTGTG	ATCGCCATCG	GGGTCGTGCA	GGCACTGATT	600
GTGGGGTACG	CATTCCACTT	CCCGCACCTG	CTGAGCCCGC	AGATCCAGCG	CTCTGCCCAC	660
AGGGCTCTGT	ACCGACGACA	CGTCCTGGGC	ATCGTCCTCC	AAGGCCCGGC	CCTGTGCTTT	720

GCAGCGGCCA	TCTTCTCTCT	CTTCTTTGTC	CCCTTGTCTT	ACCTGCTGAT	GGTGACTGTC	780
ATCCTCCTCC	CCTATGTCAG	CAAGGTCACC	GGCTGGTGCA	GAGACAGGCT	CCTGGGCCAC	840
AGGGAGCCCT	CGGCTCACCC	AGTGGAAGTC	TTCTCGTTTG	ACCTCCACGA	GCCACTCAGC	900
AAGGAGCGCG	TGGAAGCCTT	CAGCGACGGA	GTCTACGCCA	TCGTGGCCAC	GCTTCTCATC	960
CTGGACATCT	GCGAAGACAA	CGTCCCGGAC	CCCAAGGATG	TGAAGGAGAG	GTTCAGCGGC	1020
AGCCTCGTGG	CCGCCCTGAG	TGCGACCGGG	CCGCGCTTCC	TGGCGTACTT	CGGCTCCTTC	1080
GCCACAGTGG	GACTGCTGTG	GTTCGCCCAC	CACTCACTCT	TCCTGCATGT	GCGCAAGGCC	1140
ACGCGGGCCA	TGGGGCTGCT	GAACACGCTC	TCGCTGGCCT	TCGTGGGTGG	CCTCCCACTA	1200
GCCTACCAGC	AGACCTCGGC	CTTCGCCCGG	CAGCCCCGCG	ATGAGCTGGA	GCGCGTGCGT	1260
GTCAGCTGCA	CCATCATCTT	CCTGGCCAGC	ATCTTCCAGC	TGGCCATGTG	GACCACGGCG	1320
CTGCTGCACC	AGGCGGAGAC	GCTGCAGCCC	TCGGTGTGGT	TTGGCGGCCG	GGAGCATGTG	1380
CTCATGTTCG	CCAAGCTGGC	GCTGTACCCC	TGTGCCAGCC	TGCTGGCCTT	CGCCTCCACC	1440
TGCCTGCTGA	GCAGGTTCAG	TGTGGGCATC	TTCCACCTCA	TGCAGATCGC	CGTGCCCTGC	1500
GCCTTCCTGT	TGCTGCGCCT	GCTCGTGGGC	CTGGCCCTGG	CCACCTGCG	GGTCCTGCGG	1560
GGCCTCGCCC	GGCCCGAACA	CCCCCCGCCA	GCCCCACGG	GCCAGGACGA	CCCACAGTCC	1620
CAGCTCCTCC	CTGCCCCCTG	CTAGCAGCCA	CAGAGCCCAC	TCCCAGCCGT	CCTCACCAGA	1680
GATGGACCAG	GGAGGACAGG	ATGCTGGGCA	GGGGAAGCCA	AGTCACGGGC	AGGCCGCAGT	1740
GGTTCTTGCG	TGGCCTGGTT	TTATTTTCAT	TGTGAAATAT	CATGCTCTTA	TTTCAGTCCT	1800
CA						1802

- (2) INFORMATION FOR SEQ ID NO: 105:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 1395 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: BRSTNOT09
    - (B) CLONE: 1664634
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 105:

GTACCTCGGC TTATTTCATA AACAGGTACT GAAGGAAGCA GAGGCATGTG GAGGACTTCC 60 CCACCTCGTG CAGCTATTTG GGCCGTGGCA TCTGAAATTT CTTATTTCAG AGTCACCCCT 120 TTGATGACCT TGGCAGTGAA CTGCAGTCAT CTGTTTAGGC CTTTCCATGG CCCACGTCAA 180

TGCCGGTATT	TCTGTTTGTT	GCACATTTGA	TTTCCTTGTT	GTTGGCATTT	AGAAGGCCCT	240
CGAGCCGCAC	TGAGGGACTG	AGCCTGGTGT	ATATGGCAGC	AAGACTGGAT	GGTGGCTTTG	300
CAGCAGTCTC	CAGAGCATTC	CATGAGATCC	GGGCTCGAAA	TCCAGCATTT	CAGCCACAAA	360
CTTTGATGGA	CTTTGGCTCA	GGTACTGGTT	CTGTCACCTG	GGCTGCTCAC	AGTATTTGGG	420
GCCAGAGCCT	ACGTGAATAT	ATGTGTGTGG	ACAGATCAGC	TGCCATGTTG	GTTTTGGCAG	480
AAAAACTACT	GACAGGTGGT	TCAGAATCTG	GGGAGCCTTA	TATTCCAGGT	GTCTTTTTCA	540
GACAGTTTCT	ACCTGTATCA	CCCAAGGTGC	AGTTTGATGT	AGTAGTGTCA	GCTTTTTCCT	600
TAAGTGACCA	GCTACTGACA	TTTATACTTT	CGTGTAATTC	AAGTCTTCTG	CATATTTTCC	660
CCTTTTGTGA	ACAGGTACTG	GTGGAGAATG	GAACAAAAGC	TGGGCACAGC	CTTCTCATGG	720
ATGCCAGGGA	TCTGGTCCTT	AAGGGAAAAG	AGAAGTCACC	TTTGGACCCT	CGACCTGGTT	780
TTGTCTTTGC	CCCGTGTCCC	CATGAACTCC	CTTGTCCCCA	GTTGACCAAC	CTGGCCTGTA	840
GCTTCTCACA	GGCGTACCAT	CCCATCCCCT	TCAGCTGGAA	CAAGAAACCA	AAGGAAGAAA	900
AGTTCTCTAT	GGTGATCCTT	GCTCGGGGGT	CTCCAGAGGA	GGCTCATCGC	TGGCCCCGTA	960
TCACTCAGCC	TGTCCTTAAA	CGGCCTCGCC	ATGTGCATTG	TCACTTGTGC	TGTCCAGATG	1020
GGCACATGCA	GCATGCTGTG	CTCACAGCCC	GCCGGCACGG	CAGGTATGGG	GGGTGTGACC	1080
AAAATCAGTG	GGATGTGGCA	GGAAGCTGCA	GCCCACGCCA	GCATCTGTTT	CCACAGGGAT	1140
TTGTATCGTT	GTGCCCGTGT	CAGCTCCTGG	GGAGATCTTT	TACCTGTGCT	TACTCCGTCT	1200
GCGTTTCCTC	CATCTACGGC	TCAGGATCCC	TCTGAGAGTT	GATGAGGATG	TGTAACAAGT	1260
ATTTTCTTCT	ATCGTGCCTG	CCAGGGCTGA	AGCTGCCTGG	TATCCAGGAG	GGGAATGCTG	1320
GTATCCCCAT	ATGTCTGTGT	TTGTTTGAGA	TTTTTAATAA	TAAATAATAA	ATTTTTGAAG	1380
AATGGAAAAA	AAAAA					1395

#### (2) INFORMATION FOR SEQ ID NO: 106:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1635 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
    (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: PROSTUT10
  - (B) CLONE: 1690990
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 106:

CCCTCTTCCT TTTGCGCACG GAAGAACAAA TCACAACAAT CACACCAG GACTGAATCC 60

ATCAGCAGAT	ACTGCCCTGT	GGGAAGGGCA	GAGGAAAGAG	AAGACAGACG	GACTGACAGA	120
CACCACAGAG	GAACAGGGGA	GTTAGCCTGG	GACCAATGGA	GGAGAAGTAC	GAACCCTGGG	180
AAAAAGACGT	GTCAGATGAG	AAAGTTCCGG	AGAGTCCGAT	GTCTCATCGC	AGGTGTTACA	240
TCATCAGGGT	TTGCCATTGG	AATACTGAGT	GGAGATGGGA	AAGAGAAAAG	TTAAGGGCTG	300
AAATGGGAGG	GGAATGGGAA	GAAAAAATGA	GAGACAAGAG	GGAAATAAGA	AAAAACAAAG	360
AGAGCACAAA	GACCAGTTTA	GGAGAAAGGA	CCAATGGGGA	CAGTGGCAGA	GTGGCGAGGT	420
AGGTGAAGGA	CTGAGGCACA	GCGTCCTGTT	GTGGAGGGAG	GAAAGGCAAG	CGTTCCGAGG	480
TGGTGAAAAG	GAAGGCCTGC	TAGGCACGGT	GGGGATGAAC	GAGGATGCCA	TGAGTCACAC	540
AAAAGACAGT	GCTGGTGAGG	CCCAGCCACA	GGAGCCTCAG	ATAACTTGGT	AAAGGCATGT	600
CTCCCATTTG	GGAACTGATG	TTCCTAAGAT	CCGCACTGAC	GCTGCTCAGC	CGGTCCATCA	660
CACAGCAAAG	GCGTGAGGAA	GGGTCACTGC	CCAGCTGGAC	TCCAGGGTGG	TCCACGCATG	720
ACAGTCACAC	CGAACCTTCA	TGAGGATGTG	AACTGTTGGC	TCCAATTTAC	CATTCCCAGC	780
AATTCCACTC	AGATATTTGT	ATACTAATGT	TCACAGCAGC	GTGAACTCCA	CAGCAGGTGG	840
AGTAATGTTC	CATTGTGTGC	ATATGCCACA	TTTTGTTTAT	CCATTCATCT	GTTGATGCAC	900
ATTTCGGTTG	TTCCCACCTT	TGGGCTATTA	TTAATAATGC	TGCTGTGAAC	ATTCCCAAGA	960
GAAATAGGAA	GACGGCTTTG	CTAAGAACTA	AAAAAGGGAT	GGACAACAAG	GGCATATACC	1020
CAGGGGCAGT	GTTCTATCAT	GACAGCTTTA	CTGAGAGCAG	AGTAGTTCTG	CTCAGAATCA	1080
GAACACTTGT	TCCCTATAGC	CCCCCTGATT	GCCCCACAAC	CACCACCGCA	TACTCCCCTT	1140
TTCCCAACCA	TGGGCAGCAG	ATTGAGCTAT	TAACAGAAGT	GTCCTTTCGC	TGGATTTCTC	1200
AACCCTTTCC	TCATCGTCCA	CATAGAGAAA	CAGTAACAGA	TTGCTACTCA	CCCAACACCC	1260
AGGTCAAGTC	CAATGCAGGT	AGGAATAACA	GCAAATCCTT	CAATTTCTTG	ATTCTGCTCT	1320
TAAAAATCTT	AACAGAGGCT	TCCAGGTTCT	GAAAATATTT	TCTGCATAAA	CGTGTGACAC	1380
TCCATCACGA	AACTCCCTTT	GGTTATCTGC	TTAAACTTAT	CGCAAATGTC	TGGAACGCTG	1440
GTGGCTTCCA	AAATCAACTC	CTGGTGCTGC	TTAATTAAGG	TCAGGGCCAC	CCGGAAGATA	1500
ATCTTCGAGC	CTTCGTTAAA	CAAACAGTCC	CAGATCCGAA	GCACTGTCTC	CACGGGCAAG	1560
ATGTCCACAA	ACAGGCAGAT	GAACCAGCGG	GACACCAGCA	GCGTCCACAG	CACACCGAGA	1620
CGCTCCATCA	GGGGG					1635

# (2) INFORMATION FOR SEQ ID NO: 107:

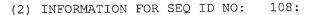
#### (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1485 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE: (A) LIBRARY: DUODNOT02 (B) CLONE: 1704050

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 107:

TTTTTGGTCC	CGNCNAAAGN	CCNAAAACCC	GGNACCCGGG	AAGCCNCCCC	AANNCNAAAN	60
TTCCCAGTTN	GAANCCCGAA	GGNAAAACCC	CGGAAAAGNA	NNCNGCCCCN	AAANTTCNCG	120
GGCNAAAACC	CGGCCNTTTT	TTCCCCCCCG	GGCGGCCGTT	TTGGGCCCCN	GANTTTCCAT	180
TTAAANTNCC	NAGNCTTGGG	CAACCTAACC	AGGNTTTTCC	CCCAANCTGG	AAAAAGCCGG	240
GCCAAGTTGA	GCCGCACCCG	CCCCAGAAGT	TCAAGGGCCC	CCGGCCTCCT	GCGCTCCTGC	300
CGCCGGGACC	CTCGACCTCC	TCAGAGCAGC	CGGCTGCCGC	CCCGGGAAGA	TGGCGAGGAG	360
GAGCCGCCAC	CGCCTCCTCC	TGCTGCTGCT	GCGCTACCTG	GTGGTCGCCC	TGGGCTATCA	420
TAAGGCCTAT	GGGTTTTCTG	CCCCAAAAGA	CCAACAAGTA	GTCACAGCAG	TAGAGTACCA	480
AGAGGCTATT	TTAGCCTGCA	AAACCCCAAA	GAAGACTGTT	TCCTCCAGAT	TAGAGTGGAA	540
GAAACTGGGT	CGGAGTGTCT	CCTTTGTCTA	CTATCAACAG	ACTCTTCAAG	GTGATTTTAA	600
AAATCGAGCT	GAGATGATAG	ATTTCAATAT	CCGGATCAAA	AATGTGACAA	GAAGTGATGC	660
GGGGAAATAT	CGTTGTGAAG	TTAGTGCCCC	ATCTGAGCAA	GGCCAAAACC	TGGAAGAGGA	720
TACAGTCACT	CTGGAAGTAT	TAGTGGCTCC	AGCAGTTCCA	TCATGTGAAG	TACCCTCTTC	780
TGCTCTGAGT	GGAACTGTGG	TAGAGCTACG	ATGTCAAGAC	AAAGAAGGGA	ATCCAGCTCC	840
TGAATACACA	TGGTTTAAGG	ATGGCATCCG	TTTGCTAGAA	AATCCCAGAC	TTGGCTCCCA	900
AAGCACCAAC	AGCTCATACA	CAATGAATAC	AAAAACTGGA	ACTCTGCAAT	TTAATACTGT	960
TTCCAAACTG	GACACTGGAG	AATATTCCTG	TGAAGCCCGC	AATTCTGTTG	GATATCGCAG	1020
GTGTCCTGGG	AAACGAATGC	AAGTAGATGA	TCTCAACATA	AGTGGCATCA	TAGCAGCCGT	1080
AGTAGTTGTG	GCCTTAGTGA	TTTCCGTTTG	TGGCCTTGGT	GTATGCTATG	CTCAGAGGAA	1140
AGGCTACTTT	TCAAAAGAAA	CCTCCTTCCA	GAAGAGTAAT	TCTTCATCTA	AAGCCACGAC	1200
AATGAGTGAA	AATGATTTCA	AGCACACAAA	ATCCTTTATA	ATTTAAAGAC	TCCACTTTAG	1260
AGATACACCA	AAGCCACCGT	TGTTACACAA	GTTATTAAAC	TATTATAAAA	CTCTGCTTTG	1320
TCCGACATTT	GCAAAGAGGT	ACACGAGGAA	ATGGAATTGG	TATTTCATTT	TAATTTTCAT	1380
GACTACTAAC	TCACCTGAAC	TTGCTATTTT	AAACAAATAG	TTCTGTCGAC	ACCTAAAATA	1440
TAATCTGGCT	TCTTGTGTCT	GGACTAAGTT	AAAAGAATTA	AAATA		1485



- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 810 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: PROSNOT16
  - (B) CLONE: 1711840
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 108:

CGAGTGAGCG	CGCGGCGGCC	CCTGGTCCGC	CCGGCCGCGG	CCGATCTAGG	GGCTGGGGGC	60
TGGAGGCGGG	GGTGGGGGTC	TGAGCTGCGT	CCTGGGCTCG	AGGCGTCCCC	CGGGGAGTCG	120
CCTCTTAGCG	GTGCGTCCGG	GCTAGCGGCG	AGGGGCCGCC	CCAAGTCTTC	CCACCGCCGC	180
CACCTTAGCA	GCCCGACTTG	GGGCCTGGAA	AGTGGAGCAC	GCGGAGGTGG	GAGGGCCCTG	240
CACGCGGCCC	CCGGTGGGGA	AGGGGACGGG	CCAGGGATTC	AGACTCGGGC	TCTCCCCTCA	300
GGATGCAGCA	CCGAGGCTTC	CTCCTCCTCA	CCCTCCTCGC	CCTGCTGGCG	CTCACCTCCG	360
CGGTCGCCAA	AAAGCAAGAT	AAGGTGAAGA	AGGGCGGCCC	GGGGAGCGAG	TGCGCTGAGT	420
GGGCCTGGGG	GCCCTGCACC	CCCAGCAGCA	AAGGATTTGC	GGCAGTGGGT	TTTCCGCGAG	480
GGCCACCTTG	GGGGGCCCA	AGAACCCAAC	CGGCAGTCCT	GGTTGAAAGG	GTTGCCCCTG	540
GAAAGTTGGA	AAGAAAGGAG	TTTTGGGCAC	CCGGACTTTG	GAAAGTTGGC	CAAATTTTTT	600
GGAAGAAAAC	TTGGCGGGTC	TGCCGGTCCG	TTAAATGGGG	GAGGGGACAA	AAGAATTGAA	660
AGCCGAAAAA	ATGCTTTCTC	CGCCGCCAAG	AGAGGTCGAA	CCCGCGTCTG	GCAAGAAGAG	720
AAAAGGGCGC	GCCCACACTG	TTAACAACAA	TATGGCGCCT	GAACAGTTGG	TGGCACCACA	780
GGGGGAGGGA	GACACATACT	TGCGCGCGGT				810

- (2) INFORMATION FOR SEQ ID NO: 109:
- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1064 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 109:

TTCCTGGGGC TCCGGGGCGC GGAGAAGCTG CATCCCAGAG GAGCGCGTCC AGGAGCGGAC

The state of the s

CCGGGAGTGT	TTCAAGAGCC	AGTGACAAGG	ACCAGGGGCC	CAAGTCCCAC	CAGCCATGCA	120
GACCTGCCCC	CTGGCATTCC	CTGGCCACGT	TTCCCAGGCC	CTTGGGACCC	TCCTGTTTTT	180
GGCTGCCTCC	TTGAGTGCTC	AGAATGAAGG	CTGGGACAGC	CCCATCTGCA	CAGAGGGGGT	240
AGTCTCTGTG	TCTTGGGGCG	AGAACACCGT	CATGTCCTGC	AACATCTCCA	ACGCCTTCTC	300
CCATGTCAAC	ATCAAGCTGC	GTGCCCACGG	GCAGGAGAGC	GCCATCTTCA	ATGAGGTGGC	360
TCCAGGCTAC	TTCTCCCGGG	ACGGCTGGCA	GCTCCAGGTT	CAGGGAGGCG	TGGCACAGCT	420
GGTGATCAAA	GGCGCCCGGG	ACTCCCATGC	TGGGCTGTAC	ATGTGGCACC	TCGTGGGACA	480
CCAGAGAAAT	AACAGACAAG	TCACGCTGGA	GGTTTCAGGT	GCAGAACCCC	AGTCCGCCCC	540
CGACACTGGG	TTCTGGCCTG	TGCCAGCGGT	GGTCACTGCT	GTCTTCATCC	TCTTGGTCGC	600
TCTGGTCATG	TTCGCCTGGT	ACAGGTGCCG	CTGTTCCCAG	CAACGCCGGG	AGAAGAAGTT	660
CTTCCTCCTA	GAACCCCAGA	TGAAGGTCGC	AGCCCTCAGA	GCGGGAGCCC	AGCAGGGCCT	720
GAGCAGAGCC	TCCGCTGAAC	TGTGGACCCC	AGACTCCGAG	CCCACCCCAA	GGCCGCTGGC	780
ACTGGTGTTC	AAACCCTCAC	CACTTGGAGC	CCTGGAGCTG	CTGTCCCCCC	AACCCTTGTT	840
TCCATATGCC	GCAGACCCAT	AGCCGCCTGC	AAGGAAGAGA	GGACACAGGA	GTAGCCACCC	900
TGAGTGCCGA	CCTTTGGTGG	CGGGGGCCTG	GGTCTCTCGT	CCCCACCCGG	AAGGGCACAA	960
GACACCGGGC	TTTGCTTGGC	AAGGCTTGGG	GCCTCTTGTG	GTCAACCCAG	TTCCCTTGGG	1020
TGCCGTTGCA	GAACCCCTTA	GCCCCTTCCA	ACGTCGACCA	GGTT		1064

#### (2) INFORMATION FOR SEQ ID NO: 110:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1031 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
    (D) TOPOLOGY: linear

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 110:

AGTTCCTGCA	GGTGCCGGCG	GTGACGCGGG	CTTACACCGC	AGCCTGTGTC	CTCATCCACC	60
GCCGCGGTGC	AGCTGGAGCT	CCTCAGCCCC	TTTCAACTCT	ACTTCAACCC	GCACCTTGTG	120
TTCCGGAAGT	TCCAGGTGAG	GCCGCCTCGC	GCCGCGCACC	TGGGGCCCGA	CCCACCCACC	180
CCGCACCTGA	CCGCCCGTCC	CCCGTAGGTC	TGGAGGCTCG	TCACCAACTT	CCTCTTCTTC	240
GGGCCCCTGG	GATTCAGCTT	CTTCTTCAAC	ATGCTCTTCG	TGTATCCTGC	GCCTGCGGAC	300
ACGGGCTGGG	TGGAGGGCAG	GCCGGCCGGG	CTGGGAGAGA	GGCCGGGACG	GGGAAACTGA	360

GGCCCCGCCT GGTGGCACT	CCTATACCGA	CGCCGTAGGT	TCCGCTACTG	CCGCATGCTG	420
GAAGAGGGCT CCTTCCGCG	G CCGCACGGCC	GACTTCGTCT	TCATGTTTCT	CTTCGGGGGC	480
GTCCTTATGA CCGTATCCT	CCCGCAGGCT	CTGGAACCTC	GGGCTAGGGC	GCCTCGGCGT	540
CCAGCCTGTG TTGGTCCTG	GGCCAACACA	GCCATGCCAG	AGAGGGACAC	AGTCGCTGTC	600
TCCAGCTTAG CACCGTTCC	T GCCTTGGGCG	CTCATGGGCT	TCTCGCTGCT	GCTGGGCAAC	660
TCCATCCTCG TGGACCTGC	T GGGGATTGCG	GTGGGCCATA	TCTACTACTT	CCTGGAGGAC	720
GTCTTCCCCA ACCAGCCTG	G AGGCAAGAGG	CTCCTGCAGA	CCCCTGGCTT	CCTAAAGCTG	780
CTCCTGGATG CCCCTGCAG	A AGACCCCAAT	TACCTGCCCC	TCCCTGAGGA	ACAGCCAGGA	840
CCCCATCTGC CACCCCCGC	A GCAGTGACCC	CCACCCAGGG	CCAGGCCTAA	GAGGCTTCTG	900
GCAGCTTCCA TCCTACCCA	T GACCCCTACT	TGGGGCAGAA	AAAACCCATC	CTAAAGGCTG	960
GGCCCATGCA AGGGCCCAC	C TGAATAAACA	GAATGAGCTG	САААААААА	AAAAAAGGGC	1020
GGCCGTCGCG A					1031

# (2) INFORMATION FOR SEQ ID NO: 111:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2316 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: PROSTUT12
- (B) CLONE: 1812375

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 111:

GCTGGATAAG	ACACCAGGGG	AGTCACTACA	TGGTTACCGC	ATCTGTATCC	AGGCCATCCT	60
GCAAGACAAG	CCCAAGATTG	CCACGGCAAA	CCTAGGCAAG	TTCCTGGAAC	TGCTGAGGTC	120
CCACCAGAGC	CGACCAGCAA	AGTGTCTCAC	CATCATGTGG	GCCCTGGGTC	AAGCAGGTTT	180
TGCCAACCTC	ACCGAGGGAC	TGAAAGTGTG	GCTGGGGATC	ATGCTGCCTG	TGCTGGGCAT	240
CAAGTCTCTG	TCTCCCTTTG	CCATCACATA	CCTGGATCGG	CTGCTCCTGA	TGCATCCCAA	300
CCTTACCAAG	GGCTTCGGCA	TGATTGGCCC	CAAGGACTTC	TTCCCACTTC	TGGACTTTGC	360
CTATATGCCG	AACAACTCCC	TGACACCCAG	CCTGCAGGAG	CAGCTGTGTC	AGCTCTACCC	420
CCGACTGAAA	ATGCTGGCAT	TTGGAGCAAA	GCCGGATTCC	ACCCTGCATA	CCTACTTCCC	480
TTCTTTCCTG	TCCAGAGCCA	CCCCTAGCTG	TCCCCTGAG	ATGAAGAAAG	AGCTCCTGAG	540
CAGCCTGACT	GAGTGCCTGA	CGGTGGACCC	CCTCAGTGCC	AGCGTCTGGA	GGCAGCTGTA	600

# PF-0459 US CCCTAAGCAC CTGTCACAGT CCAGCCTTCT GCTGGAGCAC TTGCTCAGCT CCTGGGAGCA 660 GATTCCCAAG AAGGTACAGA AGTCTTTGCA AGAAACCATT CAGTCCCTCA AGCTTACCAA 720 CCAGGAGCTG CTGAGGAAGG GTAGCAGTAA CAACCAGGAT GTCGTCACCT GTGACATGGC 780 CTGCAAGGGC CTGTTGCAGC AGGTTCAGGG TCCTCGGCTG CCCTGGACGC GGCTCCTCCT 840 AGCGTGTGCC AAGCTCTACT CCTACAGTCT GCAAGGCTAC AGCTGGCTGG GGGAGACACT 1020 GCCGCTCTGG GGCTCCCACC TGCTCACCGT GGTGCGGCCC AGCTTGCAGC TGGCCTGGGC 1080 TCACACCAAT GCCACAGTCA GCTTCCTTTC TGCCCACTGT GCCTCTCACC TTGCGTGGTT 1140 TGGTGACAGT CTCACCAGTC TCTCTCAGAG GCTACAGATC CAGCTCCCCG ATTCCGTGAA 1200 TCAGCTACTC CGCTATCTGA GAGAGCTGCC CCTGCTTTTC CACCAGAATG TGCTGCTGCC 1260 ACTGTGGCAC CTCTTGCTTG AGGCCCTGGC CTGGGCCCAG GAGCACTGCC ATGAGGCATG 1320 CAGAGGTGAG GTGACCTGGG ACTGCATGAA GACACAGCTC AGTGAGGCTG TCCACTGGAC 1380 CTGGCTTTGC CTACAGGACA TTACAGTGGC TTTCTTGGAC TGGGCACTTG CCCTGATATC 1440 CCAGCAGTAG GCCCTGCCTT CCTGGCCACT GATTTCTGCA TGGGTAGACC ATCCAAGACT 1500

GTTGCTGCTG GTCTTCGCTG TAGGCTTCCT GTGCCATGAC CTCCGGTCAC ACAGCTCCTT 900 CCAGGCCTCC CTTACTGGCC GGTTGCTTCG ATCATCTGGC TTCTTACCTG CTAGCCAACA 960 GCAGCGGGTA GAAGGTGGCA GTTCTTCATG GGAGTCTTTT TAACTTGGTG CCTGAGTTCT 1560 CTCCTAGGCA AGTGGCCAGT TGCCTCCACC TCAGTTCTTC CATCTTTGGT GGGGACAGGG 1620 CCCAGCAGCA TCTCAGCCTC CTACCCACAA TTCCACTGAA CACTTTTCTG GCCCTACTGC 1680 ACATGGCCCC CAGCCTCCAT CCTTGTGCTG GTAGCCTCTC ACAACTCCGC CCTTGCCCTC 1740 TGCCTTCCAC TTCCTTCCAT CTCATTTCTA AACCCCAAAC AGCTCATCTC TAAAAAGATA 1800 GAACTCCCAG CAGGTGGCTT CTGTGTTCTT CTGACAAATG ATTCCTGCTT CTCCAGACTT 1860 TAGCAGCCTC CTGTTCCCAT TCTTGGTCAC AGCTCTAGCC ACAGCAGAAG GAAAGGGGCT 1920 TCCAGAAGAA TATAGCACCG CATTGGGAAA CAGCAGCCTC ACCTCCACCT GAAGCCTGGG 1980 TGTGGCTGTC AGTGGACATG GGGAGCTGGA TGGAAATGCC TCTCACTTCA AAATGCCCAG 2040 CCTGCCCCAA ATGCCTCTAA GCCCCTCCCT GTCCCCTCCC TTGTAGTCCT ACTTCTTCCA 2100 ACTITICATT CCCCATCATG CTGGGGGTCT TGGTCACAAG GCTCAGCTTC TCTCCACTGT 2160 CCATCCCTCC TATCATCTGT AGAGCAGAGC ACAGGCAGTT GTGTGCCTTG GGCCCAGGGA 2220 ACCCTCCATC AACCTGAGAC AGGACTCAGT ATATGGTTCT TGGGTATGCC CTACCAGGTG 2280 GAATAAAGGA CACAGATTTG AAAAAAAAAA AAAAAA 2316

- (2) INFORMATION FOR SEQ ID NO: 112:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 1169 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: PROSNOT20 (B) CLONE: 1818761
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 112:

• •	-					
AGCAAGGAGC	CAGAGGCCAT	GCAGTGGCTC	AGGGTCCGTG	AGTCGCCTGG	GGAGGCCACA	60
GGACACAGGG	TCACCATGGG	GACAGCCGCC	CTGGGTCCCG	TCTGGGCAGC	GCTCCTGCTC	120
TTTCTCCTGA	TGTGTGAGAT	CCCTATGGTG	GAGCTCACCT	TTGACAGAGC	TGTGGCCAGC	180
GGCTGCCAAC	GGTGCTGTGA	CTCTGAGGAC	CCCCTGGATC	CTGCCCATGT	ATCCTCAGCC	240
TCTTCCTCCG	GCCGCCCCA	CGCCCTGCCT	GAGATCAGAC	CCTACATTAA	TATCACCATC	300
CTGAAGGGTG	ACAAAGGGGA	CCCAGGCCCA	ATGGGCCTGC	CAGGGTACAT	GGGCAGGGAG	360
GGTCCCCAAG	GGGAGCCTGG	CCCTCAGGGC	AGCAAGGGTG	ACAAGGGGGA	GATGGGCAGC	420
CCCGGCGCCC	CGTGCCAGAA	GCGCTTCTTC	GCCTTCTCAG	TGGGCCGCAA	GACGGCCCTG	480
CACAGCGGCG	AGGACTTCCA	GACGCTGCTC	TTCGAAAGGG	TCTTTGTGAA	CCTTGATGGG	540
TGCTTTGACA	TGGCGACCGG	CCAGTTTGCT	GCTCCCCTGC	GTGGCATCTA	CTTCTTCAGC	600
CTCAATGTGC	ACAGCTGGAA	TTACAAGGAG	ACGTACGTGC	ACATTATGCA	TAACCAGAAA	660
GAGGCTGTCA	TCCTGTACGC	GCAGCCCAGC	GAGCGCAGCA	TCATGCAGAG	CCAGAGTGTG	720
ATGCTGGACC	TGGCCTACGG	GGACCGCGTC	TGGGTGCGGC	TCTTCAAGCG	CCAGCGCGAG	780
AACGCCATCT	ACAGCAACGA	CTTCGACACC	TACATCACCT	TCAGCGGCCA	CCTCATCAAG	840
GCCGAGGACG	ACTGAGGGCC	TCTGGGCCAC	CCTCCCGGCT	GGAGAGCTCA	GGTGCTGGTC	900 .
CCGTCCCCTG	CAGGGCTCAG	TTTGCACTGC	TGTGAAGCAG	GAAGGCCAGG	GAGGTCCCCG	960
GGGACCTGGC	ATTCTGGGGA	GACCCTGCTT	CTATCTTGGC	TGCCATCATC	CCTCCCAGCC	1020
TATTTCTGCT	CCTCTCTTCT	CTCTTGGACC	TATTTTAAGA	AGCTTGCTAA	CCTAAATATT	1080
CTAGAACTTT	CCCAGCCTCG	TAGCCCAGCA	CTTCTCAAAC	TTGGAAATGC	ATGCGAATCA	1140
CCCGGGGTTC	GTGTTAAATG	CAGATTCTG				1169

- (2) INFORMATION FOR SEQ ID NO: 113:
  - (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1530 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: single(D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: GBLATUT01
- (B) CLONE: 1824469
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 113:

TCACAGACTG CGGAG	GTGGGT CAGGGGCTC	GC GAGGGCTGCC	CCAAGTCCTA	CCGGGTTTGC	60
ACGGGCGCGC CCGGC	CTCCGC CCGCAAGT(	GC GCCTTCCTGA	CTTACTGCTG	GGTGCGCGGG	120
GCTGGGGGTG CGAGT	PACCAC CCCTGAAG	TC TCTTCCTGGG	CGACCTCCGG	GGCCTCATTC	180
TAGGCCTCCT TAAAG	GAGAAG GATCTAAA	TT AGGAAAAGGA	AGTGCCCTTA	TCCACGACCA	240
AGCTCTTCCA CCTGC	CGGAGC TCGCTTAG	TC TGCACCTCAA	CCGTGCGGAA	AGTGACTGCC	300
CTGTTTACTG AGGAA	AAAACT GGGGCTCA	GA AAGATACCAT	GAGTAGTTTG	AAACAGGAAC	360
AAAATCTTCT GAAAG	GCTCGG AGCAGAAG	CC TTTTTGGTCA	ACATGGAGGA	AAAAAGACGG	420
CGAGCCCGAG TTCAG	GGGAGC CTGGGCTG	CC CCTGTTAAAA	GCCAGGCCAT	TGCTCAGCCA	480
GCTACCACTG CTAAC	GAGCCA TCTCCACC.	AG AAGCCTGGCC	C AGACCTGGAA	GAACAAAGAG	540
CATCATCTCT CTGAC	CAGAGA GTTTGTGT	TC AAAGAACCTO	C AGCAGGTAGT	ACGTAGAGCT	600
CCTGAGCCAC GAGTO	GATTGA CAGAGAGG	GT GTGTATGAAA	A TCAGCCTGTC	ACCCACAGGT	660
GTATCTAGGG TCTG	TTTGTA TCCTGGCT	TT GTTGACGTGA	A AAGAAGCTGA	CTGGATATTG	720
GAACAGCTTT GTCAA	AGATGT TCCCTGGA	AA CAGAGGACCO	G GCATCAGAGA	GGATATAACT	780
TATCAACAAC CAAGA	ACTTAC AGCATGGT	AT GGAGAACTT	C CTTACACTTA	TTCAAGAATC	840
ACTATGGAAC CAAA	TCCTCA CTGGCACC	CT GTGCTGCGC	A CACTAAAGAA	CCGCATTGAA	900
GAGAACACTG GCCA	CACCTT CAACTCCT	TA CTCTGCAAT	C TTTATCGCAA	TGAGAAGGAC	960
AGCGTGGACT GGCA	CAGTGA TGATGAAC	CC TCACTAGGG	A GGTGCCCCAT	TATTGCTTCA	1020
CTAAGTTTTG GTGC	CACACG CACATTIG	AG ATGAGAAAG	A AGCCACCACC	AGAAGAGAAT	1080
GGAGACTACA CATA	TGTGGA AAGAGTGA	AG ATACCCTTG	G ATCATGGTAC	CTTGTTAATC	1140
ATGGAAGGAG CGAC	CACAAGC TGACTGGC	AG CATCGAGTG	C CCAAAGAATA	CCACTCTAGA	1200
GAACCGAGAG TGAA	ACCTGAC CTTTCGG	CA GTCTATCCA	G ACCCTCGAGG	GGCACCCTGG	1260
TGACGTCAGA GCTT	TGAGAG AGAAGCTI	CA CTGAAACGG	A GCAAACCTTC	CACTGAGAAG	1320
CCACTTCAAG AGGC	CTGGTGC TGCTAGAT	CT CATGATGTG	G CTGTTGGGAA	A GATGGTGGGG	1380
TTTGTTTGCC AGCT	TTGGAGT CCTATTA	AAT GAAAGCCAG	C AACTCATGT	GGTAATAGGT	1440
CTACTGTGGG AACA	AGTTATC CCTAACCA	ACA GCTCAAAAT	C GCTATCATC	TTAGGCAAAT	1500

#### (2) INFORMATION FOR SEQ ID NO: 114:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1336 base pairs

  - (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: PROSNOT19
  - (B) CLONE: 1864292
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 114:

AGCTCGTACC	CCTCGAGTGA	AATTCTGAAA	TGAAGATGGA	GGAGGCAGTG	GGAAAAGTTG	60
AAGAACTCAT	TGAGTCCGAA	GCCCCACCAA	AAGCATCTGA	ACAAGAGACA	GCCAAGGAGG	120
AAGATGGATC	TGTAGAACTG	GAATCTCAAG	TTCAGAAAGA	TGGTGTAGCG	GATTCTACAG	180
TTATTTCTTC	AATGCCCTGC	TTGTTGATGG	AACTGAGAAG	GGACTCTTCT	GAGTCTCAGT	240
TAGCATCCAC	AGAGAGTGAC	AAGCCTACAA	CTGGCCGAGT	TTATGAGAGT	GACCCCTCTA	300
ATCACTGCAT	GCTTTCCCCT	TCCTCTAGTG	GTCACCTGGC	TGATTCAGAT	ACGTTGTCTT	360
CCGCAGAAGA	GAATGAACCC	TCTCAGGCAG	AAACGGCGGT	AGAAGGAGAC	CCTTCAGGAG	420
TGTCTGGTGC	CACAGTTGGG	CGCAAGTCTA	GGCGGTCCCG	ATCTGAAAGT	GAAACTTCCA	480
CTATGGCTGC	CAAGAAAAAC	CGGCAATCCA	GTGATAAACA	GAATGGCCGA	GTCGCCAAGG	540
TTAAAGGTCA	TCGGAGCCAA	AAGCACAAGG	AGAGGATCAG	GCTACTGAGG	CAGAAACGGG	600
AGGCTGCTGC	AAGGAAGAAA	TATAACCTGC	TGCAGGACAG	TAGTACCAGT	GATAGTGACC	660
TGACTTGTGA	CTCAAGCACG	AGCTCATCAG	ATGATGATGA	AGAGGTTTCA	GGGAGCAGCA	720
AGACAATCAC	TGCAGAGATA	CCAGATGGAC	CTCCAGTTGT	AGCTCATTAT	GATATGTCTG	780
ACACCAACTC	TGACCCAGAA	GTGGTAAATG	TGGACAATTT	ATTGGCGGCT	GCAGTAGTTC	840
AAGAGCACAG	TAATTCTGTA	GGCGGCCAGG	ACACAGGAGC	TACCTGGAGG	ACCAGCGGGC	900
TTCTAGAGGA	GCTGAATGCA	GAGGCAGGTC	ATTTGGATCC	AGGATTCCTA	GCAAGTGACA	960
AAACATCTGC	TGGCAATGCG	CCACTCAATG	AAGAAATTAA	CATTGCGTCT	TCAGATAGTG	1020
AAGTAGAGAT	TGTGGGAGTT	CAGGAACATG	CAAGGTGTGT	TCATCCTCGA	GGTGGTGTGA	1080
TTCAGAGTGT	TTCTTCATGG	AAGCATGGCT	CGGGCACGCA	GTATGTTAGC	ACCAGGCAAA	1140
CACAGTCATG	GACTGCTGTG	ACTCCCCAGC	AGACTTGGGC	TTCACCAGCA	GAAGTTGTTG	1200
ACCTTACCTT	GGATGAGGAT	AGCAGGCGTA	AATACCTACT	GTAATACAAT	GTCACTGTGT	1260

TTCCTCTGCA CTGTTCCCTT CCACTTCCTC ATCCTCTTTG TGACATGGAA GTTCATTGTC 1320 ATAGGGGTAC GGAGCT 1336

- (2) INFORMATION FOR SEQ ID NO:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 1742 base pairs
      (B) TYPE: nucleic acid
      (C) STRANDEDNESS: single

    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: THP1NOT01
    - (B) CLONE: 1866437
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 115 :

GCCCGCCCC	CTCCCCGCCC	GCCTTCCCGG	TGACCTTCAG	GGGCCCGGGT	GGCGGGCGCA	60
GGCCCCTGCG	GCGGCGGCGG	GATGTTCGTG	CAGGAGGAGA	AGATCTTCGC	GGGCAAGGTG	120
CTGCGGCTGC	ACATCTGCGC	GTCCGACGGC	GCCGAGTGGC	TGGAGGAGGC	CACCGAGGAC	180
ACCTCGGTGG	AGAAGCTCAA	GGAGCGCTGC	CTCAAGCACT	GTGCTCATGG	GAGCTTAGAA	240
GATCCCAAAA	GTATAACCCA	TCATAAATTA	ATCCACGCTG	CCTCAGAGAG	GGTGCTGAGT	300
GATGCCAGGA	CCATCCTGGA	AGAGAACATC	CAGGACCAAG	ATGTCCTATT	ATTGAAAAAA	360
AAGCGTGCTC	CATCACCACT	TCCCAAGATG	GCTGATGTCT	CAGCAGAAGA	AAAGAAAAAA	420
CAAGACCAGA	AAGCTCCAGA	TAAAGAGGCC	ATACTGCGGG	CCACCGCCAA	CCTGCCCTCC	480
TACAACATGG	ACCGGGCCGC	GGTCCAGACC	AACATGAGAG	ACTTCCAGAC	AGAACTCCGG	540
AAGATACTGG	TGTCTCTCAT	CGAGGTGGCG	CAGAAGCTGT	TAGCGCTGAA	CCCAGATGCG	600
GTGGAATTGT	TTAAGAAGGC	GAATGCAATG	CTGGACGAGG	ACGAGGATGA	GCGTGTGGAC	660
GAGGCTGCCC	TGCGGCAGCT	CACGGAGATG	GGCTTTCCGG	AGAACAGAGC	CACCAAGGCC	720
CTTCAGCTGA	ACCACATGTC	GGTGCCTCAG	GCCATGGAGT	GGCTAATTGA	ACACGCAGAA	780
GACCCGACCA	TAGACACGCC	TCTTCCTGGC	CAAGCTCCCC	CAGAGGCCGA	GGGGCCACA	840
GCAGCTGCCT	CCGAGGCTGC	CGCGGGAGCC	AGCGCCACCG	ATGAGGAGGC	CAGAGATGAG	900
CTGACGGAAA	TCTTCAAGAA	GATCCGGAGG	AAAAGGGAGT	TTCGGGCTGA	TGCTCGGGCC	960
GTCATTTCCC	TGATGGAGAT	GGGGTTCGAC	GAGAAAGAGG	TGATAGATGC	CCTCAGAGTG	1020
AACAACAACC	AGCAGAATGC	CGCGTGCGAG	TGGCTGCTGG	GGGACCGGAA	GCCCTCTCCG	1080
GAGGAGCTGG	ACAAGGGCAT	CGACCCCGAC	AGTCCTCTCT	TTCAGGCCAT	CCTGGATAAC	1140
CCGGTGGTGC	AGCTGGGCCT	GACCAACCCG	AAAACATTGC	TAGCATTTGA	AGACATGCTG	1200

GAGAACCCAC TGAACAGCAC CCAGTGGATG AATGATCCAG AAACGGGGCC TGTCATGCTG 1260
CAGATCTCTA GAATCTCCA GACACTAAAT CGCACGTAGG TGGCGTTGTT CCACTCGGCT 1320
ATCAGGCCAC AGCAGCCCC TGGTGCGCC CGAGACCGGG CAGAGTGGAC CTCACCTGGA 1380
AACTCACCTT CAGCGCCTCA GCCCTGGACT GTTAGAGGTG CTGCAGCTGC TCCTGCTCT 1440
TGATCTTATT GCTTATAAAC TTTGGTGACG GTAGTGTTA AGGCCGTATT TTTAGCATCT 1500
AGTGCTCCTC TGGGCTCTTG AGTTGCTCG TGAATTGCCG TGTAGACATT TGCTTGGAGA 1620
GTCCACTTGT TATTTGACGG AGGTAGGTTT CAACCCAGAG TTAATGTCAA GCATGCTAAT 1680

TTAACTAGTC ACTCACAGAT GACTTTTCTT TAATAAAGTC CCTTTTCCTA TTAAAAAAAA 1740

AA

#### (2) INFORMATION FOR SEQ ID NO: 116:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1074 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: SKINBIT01
  - (B) CLONE: 1871375
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 116:

GCGGTGCAGA	GGAAGCACAA	CCTCTACCGG	GACAGCATGG	TCATGCACAA	CAGCGACCCC	60
AACCTGCACC	TGCTGGCCGA	GGGCGCCCC	ATCGACTGGG	GCGAGGAGTA	CAGCAACAGC	120
GGCGGGGGCG	GCAGCCCAGC	CCCAGCACCC	CGGAGTCAGC	CACCCTCTCG	GAAAAGCGAC	180
GGCGCGCCAA	GCAGGTGGTC	TCTGTGGTCC	AGGATGAGGA	GGTGGGGCTG	CCCTTTGAGG	240
CTAGCCCTGA	GTCACCACCA	CCTGCGTCCC	CGGACGGTGT	CACTGAGATC	CGAGGCCTGC	300
TGGCCCAAGG	TCTGCGGCCT	GAGAGCCCCC	CACCAGCCGG	CCCCTGCTC	AACGGGGCCC	360
CCGCTGGGGA	GAGTCCCCAG	CCTAAGGCCG	CCCCGAGGC	CTCCTCGCCG	CCTGCCTCAC	420
CCCTCCAGCA	TCTCCTGCCT	GGAAAGGCTG	TGGACCTTGG	GCCCCCAAG	CCCAGCGACC	480
AGGAGACTGG	AGAGCAGGTG	TCCAGCCCCA	GCAGCCACCC	CGCCCTCCAC	ACCACCACCG	540
AGGACNANTT	TCAAGGGGTG	CAAGAATTGA	AGNTTCNTAA	GGGCCAANTT	GGGGGTCCCC	600
TTGACTTGGN	TTGGNAANAT	TGGGGCAAAA	AGGGCCGGTT	TTCCCCNTTT	CCCGGGANAC	660
CCCAAGGGAA	AGGGGNTTCA	AAGCTTCTTN	GGGGGGAAA	GGGGGAANCC	CTTGGGTNTT	720

TTGTTGGCCN TTTGTGANCA NCAGCGAGGA GAGTGCAAAG GTGCAGAGTN AGTTNTAGGN 780 CANTGGGTCC CTGACTGCTG CANATGGTAA GGNCGTTNNC TTGTGGACCC AAGGCAGGNA 840 AAGNTGTGGG GAGGGAAGCT GGTNTGTGCN TTGTGGGTGG AAGCGGGGAN GGCTGTGTTG 900 NANGGCAGGG AGAGGGCNAA NTGAGTTATT TATTGGGGTT CANGTGAAAA GTTTCTTGNN 960 CCCTGTNTTG TGTTNCTGTG GGATTGATTN TAAGATNGNN AGGGGTNGGT TTTTGGGGTT 1020 TTCCTGGTTG GTGGCCAAAN GGGTTGGAAA ATNGNTGGGG GGGGNTTGGA NAAT

#### (2) INFORMATION FOR SEQ ID NO: 117:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1454 base pairs

  - (B) TYPE: nucleic acid(C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LEUKNOT03
  - (B) CLONE: 1880830
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 117 :

CCCGGGGGAG	GCCTGACCCC	CTCCGCACCA	CCGTACGGAG	CCGCATTTCC	CCCGTTTCCC	60
GAGGGGCATC	CAGCCGTGTT	GCCTGGGGAG	GACCCACCCC	CCTATTCACC	CTTAACTAGC	120
CCGGACAGTG	GGAGTGCCCC	TATGATCACC	TGCCGAGTCT	GCCAATCTCT	CATCAACGTG	180
GAAGGCAAGA	TGCATCAGCA	TGTAGTCAAA	TGTGGTGTCT	GCAATGAAGC	CACCCCAATC	240
AAGAATGCAC	CCCCAGGGAA	AAAATATGTT	CGATGCCCCT	GTAACTGTCT	CCTTATCTGC	300
AAAGTGACAT	CCCAACGGAT	TGCATGCCCT	CGGCCCTACT	GCAAAAGAAT	CATCAACCTG	360
GGGCCTGTGC	ATCCCGGACC	TCTGAGTCCA	GAACCCCAAC	CCATGGGTGT	CAGGGTTATC	420
TGTGGACATT	GCAAGAATAC	TTTTCTGTGG	ACAGAGTTCA	CAGACCGCAC	TTTGGCACGT	480
TGTCCTCACT	GCAGGAAAGT	GTCATCTATT	GGGCGCAGAT	ACCCACGTAA	GAGATGTATC	540
TGCTGCTTCT	TGCTTGGCTT	GCTTTTGGCA	GTCACTGCCA	CTGGCCTTGC	CTTTGGCACA	600
TGGAAGCATG	CACGGCGATA	TGGAGGCATC	TATGCAGCCT	GGGCATTTGT	CATCCTGTTG	660
GCTGTGCTGT	GTTTGGGCCG	GGCTCTTTAT	TGGGCCTGTA	TGAAGGTCAG	CCACCCTGTC	720
CAGAACTTCT	CCTGAGCCTG	ATGACCCACA	GACTGTGCCT	GGCCCCTCCC	TGGTGGGGAC	780
AGTGACACTA	CGAAGGGAGC	TGGGGTAGTT	AAAGGCTCCC	GGGGCTTCTA	GAAGGAAGCC	840
AAGCAGCTGC	CTTCCTTTTC	CCTGGGGAGA	GGTAGGAAGG	AACCAGGCCC	TCACTTAGGT	900
TTGGAGGGGC	AGATAAGAGC	ACTGCTGACC	ATCTGCTTTC	CTCCAAGGGT	TGCTGTGTCT	960

AGGGTGAAGT AGGCAAAACG TTGCCCTTAA AACTGGGCCC TGAAGACGGT TCCAGCCTTG 1020
TCCTTCCTGT GTGCTCCTG AGAGCCATTC CTGTCCCTTA CACATTCCAG GGCAGGGTGG 1080
GGGTGGGTAG CCCTGGGGGT TCCCCTCCCT CTTGTGCACC ATTAGGACTT TGCTGCTGCT 1140
ATTGCACTTC ACCAGAGGTT GGCTCTGGCC TCAGTACCCT CAGTCTCCTC TCCCCACATT 1200
GTGTCCTGTG GGGGTGGGGT CAGCCGCTGC TCTGTACAGA ACCACAGGAA CTGATGTGTA 1260
TATAACTATT TAATGTGGGA TATGTTCCCC TATTCCTGTA TTTCCCTTAA TTCCTCCCC 1320
CGACCTTTTT TACCCCCCCA GTTGCAGTAT TTAACTGGGC TGGGTAGGGT TGCTCAGTCT 1380
TTGGGGGAGG TTAGGGACTT ATCCTGTGCT TGTAAATAAA TAAGGTCATG ACTCTAAAAA 1440
AAAAAAAAAGG GCGG

- (2) INFORMATION FOR SEQ ID NO: 118:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 2071 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: OVARNOT07
    - (B) CLONE: 1905325
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 118:

AGCTTTGAAT TCCTGTATCT GAGAACGGAT CGTTCGAGGT GGTGGAGGGG GTTGGAATTG 60
GGGACCTACG GAAGGCTCAG CTCTTGCCAG GCCAAATTGA GACATGTCTG ACACAAGCGA 120
GAGTGGTGCA GGTCTAACTC GCTTCCAGGC TGAAGCTTCA GAAAAGGACA GTAGCTCGAT 180
GATGCAGACT CTGTTGACAG TGACCCAGAA TGTGGAGGTC CCAGAGACAC CGAAGGCCTC 240
AAAGGCACTG GAGGTCTCAG AGGATGTGAA GGTCTCAAAA GCCTCTGGGG TCTCAAAAGGC 300
CACAGAGGTC TCAAAGACCC CAGAGGCTCG GGAGGCACCT GCCACCCAGG CCTCGTCTAC 360
TACTCAGCTG ACTGATACCC AGGTTCTGGC AGCTGAAAAC AAGAGTCTAG CAGCTGACAC 420
CAAGAAACAG AATGCTGACC CGCAGGCTGT GACAATGCCT GCCACCTAGAG CCAAAAAAGGT 480
CAGCCATGTG GCTGATACAA AGGTCAATAC AAAGGCTCAG GAGACTGAGG CTGCACCCTC 540
TCAGGCCCCA GCAGATGAAC CTGAGCCTGA GAGTGCAGC GCCAGTCTC AGGAGAATCA 600
GGATACTCGG CCCAAGGTCA AAGCCAAGAA AGCCCGAAAG GTGAAGCATC TGGATGGGGA 660
AGAGGGATGGC AGCAGTGATC AGAGTCAGGC TTCTGGAACC ACAGGTGGCC GAAGGGTCTC 720
AAAAGGCTCTA ATGGCCTCAA TGGCCCGCAG GTTTCAAGGG GTCCCATAGC CTTTTGGGCC 780

CGCAGGATTC	AAGGACTCGG	TTGGCTGCTT	GGGCCCGGAG	AGCCTTGCTC	TCCCTGAGAT	840
CACCTAAAGC	CCGTAGGGCA	AGGCTCGCCG	TAGAGCTGCC	AAGCTCCAGT	CATCCCAAGA	900
GCCTGAAGCA	CCACCACCTC	GGGATGTGGC	CCTTTTGCAA	GGGAGGGCAA	ATGATTTGGT	960
GAAGTACCTT	TTGGCTAAAG	ACCAGACGAA	GATTCCCATC	AAGCGCTCGG	ACATGCTGAA	1020
GGACATCATC	AAAGAATACA	CTGATGTGTA	CCCCGAAATC	ATTGAACGAG	CAGGCTATTC	1080
CTTGGAGAAG	GTATTTGGGA	TTCAATTGAA	GGAAATTGAT	AAGAATGACC	ACTTGTACAT	1140
TCTTCTCAGC	ACCTTAGAGC	CCACTGATGC	AGGCATACTG	GGAACGACTA	AGGACTCACC	1200
CAAGCTGGGT	CTGCTCATGG	TGCTTCTTAG	CATCATCTTC	ATGAATGGAA	ATCGGTCCAG	1260
TGAGGCTGTC	ATCTGGGAGG	TGCTGCGCAA	GTTGGGGCTG	CGCCCTGGGA	TACATCATTC	1320
ACTCTTTGGG	GACGTGAAGA	AGCTCATCAC	TGATGAGTTT	GTGAAGCAGA	AGTACCTGGA	1380
CTATGCCAGA	GTCCCCAATA	GCAATCCCCC	TGAATATGAG	TTCTTCTGGG	GCCTGCGCTC	1440
TTACTATGAG	ACCAGCAAGA	TGAAAGTCCT	CAAGTTTGCC	TGCAAGGTAC	AAAAGAAGGA	1500
TCCCAAGGAA	TGGGCAGCTC	AGTACCGAGA	GGCGATGGAA	GCAGATTTGA	AGGCTGCAGC	1560
TGAGGCTGCA	GCTGAAGCCA	AGGCTAGGGC	CGAGATTAGA	GCTCGAATGG	GCATTGGGCT	1620
CGGCTCGGAG	AATGCTGCCG	GGCCCTGCAA	CTGGGACGAA	GCTGATATCG	GACCCTGGGC	1680
CAAAGCCCGG	ATCCAGGCGG	GAGCAGAAGC	TAAAGCCAAA	GCCCAAGAGA	GTGGCAGTGC	1740
CAGCACTGGT	GCCAGTACCA	GTACCAATAA	CAGTGCCAGT	GCCAGTGCCA	GCACCAGTGG	1800
TGGCTTCAGT	GCTGGTGCCA	GCCTGACCGC	CACTCTCACA	TTTGGGCTCI	TCGCTGGCCT	1860
TGGTGGAGCT	GGTGCCAGCA	CCAGTGGCAG	CTCTGGTGCC	: TGTGGTTTCT	CCTACAAGTG	1920
AGATTTTAGA	TATTGTTAAT	CCTGCCAGTC	TTTCTCTTCA	AGCCAGGGTG	CATCCTCAGA	1980
AACCTACTCA	ACACAGCACI	CTAGGCAGCC	CACTATCAATC	AATTGAAGT1	GACACTCTGC	2040
ATTAAATCTA	A TTTGCCATTT	CAAAAAAAAA	A A			2071

# (2) INFORMATION FOR SEQ ID NO: 119:

- (i) SEQUENCE CHARACTERISTICS:
  (A) LENGTH: 1236 base pairs

  - (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: BRSTTUT01
  - (B) CLONE: 1919931
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 119:

ACCTGGGACC	CCCAGAACGG	CCGCCCCTTT	TTTTTTTTT	TTTTTTTTT	TTTTTTTTT	60
TTTTTTTTT	TTTTTTTTT	TTTTTTTTTT	TTTTTTTAG	AAGGTTGAAA	CCAGGCTTAT	120
TTATTTTCAT	CTTCTTTCTG	CCATCTTTTA	ACCAACCTTC	TCAGAATAAA	ATGTGATTTT	180
rgagacagaa	TGAAACACAT	ATCCAAATTT	TAATACAGTA	AGAATAGGTA	TCCTGAATAA	240
ATGAGAACTC	TAGAAAATCA	AGGTTTCAAA	ATTCTACCCT	TCCTGGGAGT	TAAAGAAGTT	300
rggcagaaac	AGAACAAATT	AATCAGCAGA	TTCATCACCT	GCCAATTTTT	TCTGTACAAT	360
TTTCTTGATT	CTGGGAGCAT	CTGGGTCCAG	GCAGATTTTC	CTCCCATCCT	TCAGTGTGGC	420
TGCTTCTTGT	TTCATCCATG	GACCCTGCAA	GAAATTGCCC	CATGTTTCTG	TTTGTGCATC	480
ACTGAGAAAG	GAAGCATGAA	GGTCGCACAG	GTCAGGCCAT	TCCATTGCCC	TCCTGGTGCC	540
GGGTTTGCCC	TCCCAATCCT	GGGGTTGCTT	CAGGGGCTTG	TCATTCTCCA	TAGTCCCCTC	600
CACATTTCTC	AGGTTTCTGC	TCAAAAGTCA	CCTTTTGGAG	GGGTCTCCAC	CTGTCACTGT	660
GTTTGTAAGA	GCTCCTTCAG	TTTCTTTCTA	GCTCATCTCA	CTCTGGTAAT	GTCTTTGATT	720
ACCACCACCA	TCTGACCTGG	TCTTATGACC	TGTTAGCTTT	CTTCATCAGA	CGTGAGCACC	780
AGGATGGCAG	GGGCCTCATC	TGTCCTGTTC	CTCCTGTGGC	CTGGGTCCTA	GCACCATGTC	840
TGGTACAGTG	TAGATGCTCA	AGGGAAGTTT	ACTTTGTAAA	ACCACTTACC	TGGGAGATGT	900
TACTGTTAGT	CTAACCTGTA	CCATTTTGTA	AACCTCCAGC	CATTTTGCAG	ACTCTGATCA	960
CAGTGAAACG	TTCCATGGGA	ACTTGGGCCA	TGAGAAACAT	CCTTCCTAAC	CACGTGACTG	1020
CAGAAACATC	CTTATCGCGT	CCTCCTGGGC	AAAGGCCCAA	CAGCCTGACT	GCAGGGACAT	1080
CCTTGCCATA	TCCTGCTGGG	CAGCAAGCTC	TACCACCCAG	ATCCCTCCCT	CCCAGTCCCA	1140
TGATTACCCC	AGCCTGTGAG	TGGCAGTTGG	TGCTGGCACT	AAGCTGGTTT	CCTCCTCCCC	1200
AGGGTTTTGC	: TGGCAATAAA	GATGTTGCTG	TTGAAG			1236

#### (2) INFORMATION FOR SEQ ID NO: 120:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1391 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
    (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRSTNOT04
- (B) CLONE: 1969426
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 120 :

GTACTGCCCA CCACCTCCCT GGGCCACCCC TCACTCAGTG CTCCGGCTCT CTCCTCCTCC 60

TCTTCGTCCT	CCTCCACTTC	ATCTCCTGTT	TTGGGCTCCC	CCTCTTACCC	TGCTTCTTCC	120
CCTGGGGCCT	CCCCCCACCA	CCGCCGTGTG	CCCCTCAGCC	CCCTGAGTTT	GCTCGCGGGC	180
CCAGCCGACG	CCAGAAGGTC	CCAACAGCAG	CTGCCCAAAC	AGTTTTCGCC	AACAATGTCA	240
CCCACCTTGT	CTTCCATCAC	TCAGGGCGTC	CCCCTGGATA	CCAGTAAACT	GTCCACTGAC	300
CAGCGGTTAC	CCCCATACCC	ATACAGCTCC	CCAAGTCTGG	TTCTGCCTAC	CCAGCCCCAC	360
ACCCCAAAGT	CTCTACAGCA	GCCAGGGCTG	CCCTCTCAGT	CTTGTTCAGT	GCAGTCCTCA	420
GGTGGGCAGC	CCCCAGGCAG	GCAGTCTCAT	TATGGGACAC	CGTACCCACC	TGGGCCCAGT	480
GGGCATGGGC	AACAGTCTTA	CCACCGGCCA	ATGAGTGACT	TCAACCTGGG	GAATCTGGAG	540
CAGTTCAGCA	TGGAGAGCCC	ATCAGCCAGC	CTGGTGCTGG	ATCCCCCTGG	CTTTTCTGAA	600
GGGCCTGGAT	TTTTAGGGGG	TGAGGGGCCA	ATGGGTGGCC	CCCAGGATCC	CCACACCTTC	660
AACCACCAGA	ACTTGACCCA	CTGTTCCCGC	CATGGCTCAG	GGCCTAACAT	CATCCTCACA	720
GGGGACTCCT	CTCCAGGTTT	CTCTAAGGAG	ATTGCAGCAG	CCCTGGCCGG	AGTGCCTGGC	780
TTTGAGGTGT	CAGCAGCTGG	ATTGGAGCTA	GGGCTTGGGC	TAGAAGATGA	GCTGCGCATG	840
GAGCCACTGG	GCCTGGAAGG	GCTAAACATG	CTGAGTGACC	CCTGTGCCCT	GCTGCCTGAT	900
CCTGCTGTGG	AGGAGTCATT	CCGCAGTGAC	CGGCTCCAAT	GAGGGCACCT	CATCACCATC	960
CCTCTTCTTG	GCCCCATCCC	CCACCACCAT	TCCTTTCCTC	CCTTCCCCCT	GGCAGGTAGA	1020
GACTCTACTC	TCTGTCCCCA	GATCCTCTTT	CTAGCATGAA	TGAAGGATGC	CAAGAATGAG	1080
AAAAAGCAAG	GGGTTTGTCC	AGGTGGCCCC	TGAATTCTGC	GCAAGGGATG	GGCCTGGGGG	1140
AACTCAAGGG	AGGGCCTAAA	GCACTTGTAA	CTTTGAACCG	TCTGTCTGGA	GGTCAGAGCC	1200
TGTTGGAAAG	CAGGGGTAGA	GGGGAGCCCT	GGAAGCAGGG	CTTTTCCGGA	TGCCTAGGGG	1260
TGGGCAGTGC	CAGCCCCTCC	TCACCACTCT	TCCCCTTGCA	GTGGAGGAGA	GAGCCAGAGT	1320
GGATACTATT	TTTTATTAAA	TATATTATTA	TATGTTAATA	AAAAAATCAT	ATCAAAAAA	1380
AAAAAAAAAG	G					1391

#### (2) INFORMATION FOR SEQ ID NO: 121:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2183 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: UCMCL5T01 (B) CLONE: 1969948

243

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 121: CTCTGTGAAC ATATGATGAG AGAAGCCAAG ATCATGCAGT ATAAGTACCT ACTGTTCAGT 60 CTTCACGCCA TAGTGAAGCT TGGAATCCCT CAGAACACTA TTTTGGTGCA GACTTTGCTG 120 AGGGTGACCC AGGAACGTAT CAATGAGTGT GATGAGATAT GCCTTTCAGT TTTGTCAACT 180 GTTTTAGAGG CAATGGAACC ATGCAAGAAT GTTCATGTTC TACGAACGGG ATTCAGAATA 240 CTAGTTGATC AGCAAGTTTG GAAAATAGAA GATGTCTTCA CATTACAAGT TGTGATGAAG 300 TGTATTGGAA AAGATGCACC GATTGCTCTT AAGAGGAAAC TGGAGATGAA AGCCTTGAGG 360 GGATTAGACA GATTTTCTGT TTTGAATAGC CAACACATGT TTGAAGTACT AGCTGCCATG 420 AATCACCGAT CTCTTATACT CCTGGATGAA TGCAGTAAGG TGGTCCTAGA TAATATCCAT 480 GGGTGTCCTT TAAGAATAAT GATCAACATA TTGCAGTCCT GCAAAGACCT CCAGTACCAT 540 AATTTGGATC TCTTCAAGGG ACTTGCAGAT TATGTGGCTG CAACTTTCGA CATCTGGAAG 600 TTCAGAAAAG TTCTTTTTAT CCTCATTTTA TTTGAAAACC TTGGCTTTCG ACCTGTTGGT 660 TTAATGGACC TGTTTATGAA GAGAATAGTA GAGGATCCTG AATCCCTAAA CATGAAAAAC 720 ATTCTATCTA TTCTTCATAC TTACTCTTCT CTCAATCATG TCTACAAATG CCAGAACAAA 780 GAACAGTTCG TGGAAGTTAT GGCTAGTGCT CTGACTGGTT ATCTTCACAC TATTTCTTCT 840 GAAAACTTAT TGGATGCAGT ATATTCATTT TGCTTGATGA ATTACTTTCC CCTGGCTCCT 900 TTTAATCAGC TTCTGCAAAA AGACATCATC AGTGAGCTGC TGACATCAGA TGACATGAAG 960 AATGCTTACA AGCTGCATAC TTTGGATACT TGTCTAAAAC TTGATGATAC TGTCTATCTG 1020 AGGGACATAG CCTTGTCACT CCCACAGCTG CCGCGGGAGC TGCCATCGTC ACATACAAAT 1080 GCAAAGGTGG CAGAGGTGCT GAGCAGCCTT CTGGGAGGTG AAGGACACTT CTCAAAGGAT 1140 GTGCACTTGC CACACAATTA TCATATTGAT TTTGAAATCA GAATGGACAC TAACAGGAAT 1200 CAAGTGCTAC CACTTTCTGA TGTGGATACA ACTTCTGCTA CAGATATTCA AAGAGTAGCT 1260 GTGCTATGTG TTTCCAGATC TGCTTATTGT TTGGGTTCAA GCCACCCCAG AGGATTCCTT 1320 GCTATGAAAA TGCGGCATTT GAATGCAATG GGTTTTCATG TGATCTTGGT CAATAACTGG 1380 GAGATGGACA AACTAGAGAT GGAAGATGCA GTCACATTTT TGAAGACTAA AATCTATTCA 1440 GTAGAAGCTC TTCCTGTTGC TGCTGTAAAT GTGCAAAGCA CACAATAAAG TGAAAATCAA 1500 CCTTTTCATA TTAGGAGACA TGCATTTGTA AAAATTAATA AAGATGACAA GTCAGTTGTC 1560 AATGGAATTG AGCTATCTGC TAAGACAAAA AATGTTACCT CAGTTCACTA TTAAAATTAA 1620 TTTTAGGAGT GGAAGAATG TTGTTACTGC CATTTAAAAA TATGCTGAGA AAATTCCAGA 1680

AGGGTTATTT TTCCAACCAC ACCTATTCCC TCTAGTGCCC AGATATTTGA TTTGTGAGCT 1740 GTACGTTTCA CCTTTTCATC TTTGATCTAC TAAAAACTGG TTTCTTAGTT GTGAGGTGTC 1800

ACAGGCAGGT TGATGTGGGT AGTAGTCCTT GTCTTTGGAA TCTGAATATT TATACTCCTG 1860
CTCTAAGCTG TTCTAAGACT TGGGGTTATG CCTTTAAATC ATTTCAAGC ATTGGCCAAA 1920
TAATAATTGG ACAAAGTTCT AAAGTTGTCA AGTGTGTAAG AATTAGTGAG GTAGCTGTTG 1980
AAAATGAGTG AGGATAGTAT TTGTATTTGT AATAAGCACT GCAGGTAGAG ATATTCATG 2040
GCTTATAATA AGAGAAACAC AGATGAGATG TAGATGGTAA GGAGTCTTAC TGTTGTGGG 2100
AAATTAAATA TATAGCTTNA ATT 2183

# (2) INFORMATION FOR SEQ ID NO: 122:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2066 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LUNGAST01
  - (B) CLONE: 1988911
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 122:

AGAACCACTG CA	GTGGAGAC	TCCATGTGCA	AAAGAAAAA	ACCAAATGTG	AGGTCATAAA	60
GACTTTCTGC CA	AGCATGTGG	GTGACATTGT	TTCTTTGCAG	ATTTTGGCTA	TGGAAAGGGG	120
AAATGTTCTA AG	GCAGAGCCC	CGTCAAGAGC	CCACGGGACA	CATTTTGGAG	ATGACAGATT	180
TGAAGATCTG GA	AAGAGGCAA	ATCCATTCTC	TTTTAGAGAG	TTTCTGAAGA	CCAAGAACCT	240
CGGCCTCTCG AA	AAGAGGATC	CGGCCAGCAG	AATTTATGCA	AAGGAAGCCT	CGAGGCATTC	300
CCTGGGACTT GA	ACCACAACT	CCCCACCCTC	CCAAACCGGC	GGGTATGGCC	TGGAGTATCA	360
GCAGCCATTT TT	CGAGGATC	CGACAGGGGC	TGGTGACCTC	CTGGATGAGG	AGGAGGATGA	420
GGACACCGGA TG	GAGTGGGG	CCTACCTGCC	GTCCGCCATC	GAGCAGACTC	ACCCCGAGAG	480
GGTCCCTGCC GG	GCACGTCGC	CCTGCAGCAC	ATACCTTTCC	TTTTTCTCCA	CCCCGTCGGA	540
GCTGGCAGGG CC	CTGAGTCTC	TGCCCTCGTG	GGCGTTGAGT	GACACTGATT	CTCGCGTGTC	600
TCCGGCCTCT CC	CGGCAGGGA	GTCCTAGCGC	AGACTTTGCG	GTTCATGGAG	AGTCTCTGGG	660
AGACAGGCAC CI	rgcggacgc	TGCAGATAAG	TTACGACGCA	CTGAAAGATG	AAAATTCTAA	720
GCTGAGAAGA AA	AGCTGAATG	AGGTTCAGAG	CTTCTCTGAA	GCTCAAACAG	AAATGGTGAG	780
GACGCTTGAG CG	GGAAGTTAG	AAGCAAAAAT	GATCAAGGAG	GAAAGCGACT	ACCACGACCT	840
GGAGTCGGTG GT	TTCAGCAGG	TGGAGCAGAA	CCTGGAGCTG	ATGACCAAAC	GGGCTGTAAA	900

GGCAGAAAAC CACGTCGTGA AACTAAAACA GGAAATCAGT TTGCTCCAGG CGCAGGTCTC 960 CAACTTCCAG CGAGAGAATG AAGCCCTGCG GTGCGGCCAG GGTGCCAGCC TGACCGTGGT 1020 GAAGCAGAAC GCCGACGTGG CCCTGCAGAA CCTCCGGGTG GTCATGAACA GTGCACAGGC 1080 TTCCATCAAG CAACTGGTTT CCGGAGCTGA GACACTGAAT CTTGTTGCCG AAATCCTTAA 1140 ATCTATAGAC AGAATTTCTG AAGTTAAAGA CGAGGAGGAA GACTCTTGAG GACCCCTGGG 1200 TGTTCTCAGC ATGAAGCTCC GTGTATACCC TGAGGTCACC ACCGCTCGAT CTAAATGTGC 1260 AGTTGTGTCC TTAAATATGC AGTCTTCACC CAGAGTAAAG TGTTGATCGC AAGAGTCCAG 1320 TGTCGTGCCC TCAGCCAGTT CTTGGCCACC ACAATGGGAG CAGCCCTGGC CGAGTTGTCT 1380 CTGTGGTTTC TATGCAGCCC TTCTTGGCGA AATTCCTGCG ATCTTATAGA TTCTAATGAG 1440 CTCTTGGAAG ACATTGTCAT AAAAGCCAGT GATTTTAAGA AAAAGAGTGG TTCTGGAATC 1500 AATGTTTTCC AGTCCCATCC CAGAACATCA GTTGTAAGAT AAGTACAATT GGTTGTCCTT 1560 GATTTCATAA GTAGAACAAA CACTAAATGT GCCTCTGAGA TGGCCACCCC GGGCAGGGAC 1620 CTGTGCCTTC CGCCGATGCT CAGGGCTCCC TCTGGCTCCC GGGTCACTCT TGTGGCCCCA 1680 GTGGGTGGTC CCTGCAGTCA TGGCCTGAGT GCGCAGGGGC CACCGCGTGG CTGCTGCT 1740 CCTCCTCGG GACCCACGGG GACCAAGGTC ACACGTTCCG TGCTGTGAAG CTGTCCAGAT 1800 GTGCCTCTTT GGCTGGGGGT TCTGGTGGAC GTTTCAAGTG GCATTTTGTA CAATGCAGGT 1860 TAGAATTCAG GAATTTCAAG TATGTGCCCG GGTCTGTCAG GTCCCAGTTG CCTTTCTGAC 1920 GGCCCCCTC AGAGGGACGG CGATGAGCAC TAAATGCTTT TTTGACTATT TTCCTATAGA 1980 TTTTTTTAA AACTTTTTT TCCTCCTGTT CCAATTGATA GCTTTCTTAT TTAATAAATT 2040 CTGTAGTTCA CCGCAAAAA AAAAAA 2066

#### (2) INFORMATION FOR SEQ ID NO: 123:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1867 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: OVARNOT03
  - (B) CLONE: 2061561
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 123:

TGGCCAGGCT GGTCTAGAAC TCCTGACTGC AAATGATCAG CCCGCCTCAG CCACCCAAAG 60
TGTTGGGATT ACAGGTGTGA GCCACTGTGC CCAGCGTGAT TTTTTTTTT TTTAAAGCAA 120

ACTTGTCCTT	TGGTTTTGCA	GAACAGGCCT	GCTCCCTCTC	ATCTAGCCCA	TCATTTCTTG	180
					TGAATATGAT	
		CATTACCTTG				300
		GAGGTCTGAG				360
		ATCTGGCCAT				420
		TCCTTCCTCT				480
					GGGCCATAAT	540
		ACTTCCTGTT				600
		TCCTTTAAGG				660
		GGTATCTGTT				720
		TACTTGATTC				780
					AACCCTTTGC	840
					TTGGACAGAG	
CCTGTTTTCA	TGTACTGCAG	GTGGGGGTGT	GCTGACATGT	TTGCTCTTGG	TTGATGGAGA	960
AGGTACAGAG	GCCAGGGAGT	GAAAATGGTT	GACAGAAGAG	GGAAGAGTTA	GGTGTCTCAT	1020
AGTCACTCAT	AGTGGGGTGG	TCAGGGGTAA	TGGCATCTCC	CCACTTTAGG	CTTCTCAAAC	1080
AGACTTTTGA	CACCTCTCAA	GTTCAGAGCT	CTGATGTGGA	AAGACAGGAG	GTGTGGGGAA	1140
GGAGGGGGAT	TTCGTGTGTT	TGCATGAGTG	TGCGCTTCAG	GCCTTGGGAG	TTGGCAAGAG	1200
GGAGGGAAGG	AAGGAGAGCA	AAATCTTCGG	AAGGTGTTTC	TTGTACCTGA	GGGATCCTGC	1260
CCTGAATCTC	CATAGTCTCC	ACTGTGAACT	GAGGAGGGGA	GGGGTGTGCT	GGGGAATAAA	1320
TCTTGTATGA	GAACAATCAA	AAATCAAACG	AATCCCACCG	ACAGACTGCT	GCTCCTAGTG	1380
ATCTGGACTC	ACCTAGGGGG	CATCTGGGCT	GGGGTTCCAN	GCTTACGTNC	GCGTGNATGN	1440
GACGNCANAG	CTCTTCGAAA	GTGTCCCNAA	ANTNCAATTC	ATTGGCGGTG	GTTTTAAAAG	1500
TTCGGGCCTG	GGAAACCCGG	GGGNTTACCC	ATTTTATCCC	NCTTNGANGG	CANATTCCCC	1560
TTTTTCCCCA	ATTTGGGGAA	ATTTNCCAAA	NGGGNCCCGT	AACGGTTGGC	CTTTTCCCAA	1620
AATTTNGGNC	GCCCTTAATT	GGGGCGATTG	TGGGACCCGC	GCCCTTTATA	GGGGGGGCT	1680
TTAAAGCGGC	GCNGGGGGTT	CTTTGGGTGA	TTACCGGCGC	GGTTGACCCC	GGGTAAAATA	1740
TTGACAAGGG	CCCTTTAGC	G CGCGGTTCCT	TGTGGGGTTI	TCCTCCCATT	TGCTTTTTCC	1800
GCAAAAGTTI	TGGCGGGGTT	TTCCCCGGAA	. AAGGTCTTAA	AAAGCGGTGT	GCCCTCTTT	1860
GAGGGGG						1867

# (2) INFORMATION FOR SEQ ID NO: 124:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1628 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: PANCNOT04
- (B) CLONE: 2084489
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 124:

CTCTGGGTCT	GTAGCAACCG	CCCAGCGTTG	AGGCGCGGCT	CATGCCCCCA	GTATCCCGGT	60
CCAGCTATTC	CGAGGACATC	GTGGGCTCTC	GGAGAAGGCG	ACGCAGCTCC	TCGGGGAGCC	120
CACCATCCCC	GCAGAGCAGA	TGTTCCTCTT	GGGATGGCTG	TTCCCGCTCT	CACTCCCGCG	180
GCCGTGAGGG	CCTCAGGCCT	CCTTGGAGTG	AGTTGGACGT	GGGCGCTCTT	TACCCCTTTA	240
GTCGCTCTGG	GTCGCGAGGG	CGGCTCCCAA	GATTCCGCAA	CTACGCCTTC	GCGTCCTCCT	300
GGTCGACCTC	GTATAGTGGA	TATCGCTACC	ATCGTCACTG	CTATGCAGAA	GAACGGCAGT	360
CAGCGGAAGA	CTACGAGAAG	GAAGAGAGCC	ATCGGCAGAG	GAGGCTGAAG	GAGAGAGAGA	420
GGATTGGGGA	ATTGGGAGCG	CCTGAAGTGT	GGGGGCCGTC	TCCAAAGTTC	CCTCAGCTAG	480
ATTCTGACGA	ACATACCCCA	GTTGAGGATG	AAGAAGAGGT	AACGCATCAG	AAAAGCAGCA	540
GTTCAGATTC	CAACTCGGAA	GAACATAGGA	AAAAGAAGAC	CAGTCGTTCA	AGAAACAAGA	600
AAAAAAGAAA	GAATAAGTCG	TCTAAAAGAA	AGCATAGGAA	ATATTCTGAT	AGTGACAGTA	660
ACTCAGAGTC	TGACACAAAT	TCTGACTCTG	ATGATGATAA	AAAGAGAGTT	AAAGCCAAGA	720
AGAAAAAGAA	GAAAAAGAAA	CACAAAACAA	AGAAAAAGAA	GAATAAGAAA	ACCAAAAAAG	780
AATCCAGTGA	CTCAAGCTGT	AAAGACTCAG	AAGAGGACTT	GTCAGAAGCT	ACCTGGATGG	840
AGCAGCCAAA	TGTGGCAGAT	ACTATGGATT	TAATAGGGCC	AGAAGCACCT	ATAATACATA	900
CCTCTCAAGA	TGAAAAACCT	TTGAAGTATG	GCCATGCTTT	GCTTCCCGGT	GAAGGTGCAG	960
CTATGGCTGA	GTATGTAAAA	GCTGGAAAGC	GAATCCCACG	AAGAGGTGAA	ATTGGGTTGA	1020
CAAGTGAAGA	GATCGGTTCT	TTTGAATGCT	CAGGTTATGT	CATGAGTGGT	AGCAGGCATC	1080
GCAGAATGGA	GGCTGTACGA	CTGCGTAAGG	AGAACCAGAT	CTACAGTGCT	GATGAGAAGA	1140
GAGCTCTTGC	ATCCTTTAAC	CAAGAAGAGA	GACGAAAGAG	AGAAAGTAAG	ATTTTAGCCA	1200
GTTTCCGAGA	GATGGTGCAC	AAAAAGACAA	AAGAGAAAGA	TGACAAGTAA	GGACTTACTT	1260
GTTGCACAGC	AGGAATTTTA	ACAACAAAAA	TTTTATGTGA	CCAAAAGTGT	TAAAAGGCTT	1320
TACAGTGCTA	CTGTACTTAC	CATATTAGTA	AGTCCCTCAG	GAAAAAGCTT	CTTTTGAGAT	1380



ATCTTTAGCA GCTTATTTT TGTTATTTA ACTTTAAAAA GTAATATGTG CACATGGTTT 1440
TAAAAATATT CAACCATTAT AGGAGGAGAG TTAGTAAAAA GTGAATCTTT CACTTTAGCC 1500
CCTGACACCT TTCCCCCAAA AATATATATT TTGGTGTCTT ATATACAGAA TATACATTCT 1560
GTGCATATAC AAGAGTATAT GTTGCAGCAT AAAGATTAAA AGCTATTAAA GTTTTTTTC 1620
GCTCGTTA

# (2) INFORMATION FOR SEQ ID NO: 125:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1200 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: SPLNFET02
- (B) CLONE: 2203226

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 125:

GTGGCGGCGG	CGAAGGATGC	ACCCGGCAGG	CTTGGCGGCG	GCGGCTGCGG	GGACGCCCCG	60
GCTGCCCTCG	AAGCGGAGGA	TCCCTGTGTC	CCAGCCGGGC	ATGGCCGACC	CCCACCAGCT	120
TTTCGATGAC	ACAAGTTCAG	CCCAGAGCCG	GGGCTATGGG	GCCCAGCGGG	CACCTGGTGG	180
CCTGAGTTAT	CCTGCAGCCT	CTCCCACGCC	CCATGCAGCC	TTCCTGGCTG	ACCCGGTGTC	240
CAACATGGCC	ATGGCCTATG	GGAGCAGCCT	GGCCGCGCAG	GGCAAGGAGC	TGGTGGATAA	300
GAACATCGAC	CGCTTCATCC	CCATCACCAA	GCTCAAGTAT	TACTTTGCTG	TGGACACCAT	360
GTATGTGGGC	AGAAAGCTGG	GCCTGCTGTT	CTTCCCCTAC	CTACACCAGG	ACTGGGAAGT	420
GCAGTACCAA	CAGGACACCC	CGGTGGCCCC	CCGCTTTGAC	GTCAATGCCC	CGGACCTCTA	480
CATTCCAGCA	ATGGCTTTCA	TCACCTACGT	TTTGGTGGCT	GGTCTTGCGC	TGGGGACCCA	540
GGATAGGTTC	TCCCCAGACC	TCCTGGGGCT	GCAAGCGAGC	TCAGCCCTGG	CCTGGCTGAC	600
CCTGGAGGTG	CTGGCCATCC	TGCTCAGCCT	CTATCTGGTC	ACTGTCAACA	CCGACCTCAC	660
CACCATCGAC	CTGGTGGCCT	TCTTGGGCTA	CAAATATGTC	GGGATGATTG	GCGGGGTCCT	720
CATGGGCCTG	CTCTTCGGGA	AGATTGGCTA	CTACCTGGTG	CTGGGCTGGT	GCTGCGTGGC	780
CATCTTTGTG	TTCATGATCC	GGACGCTGCG	GCTGAAGATC	TTGGCAGACG	CAGCAGCTGA	840
GGGGGTCCCG	GTGCGTGGGG	CCCGGAACCA	GCTGCGCATG	TACCTGACCA	TGGCGGTGGC	900
GGCGGCGCAG	CCTATGCTCA	TGTACTGGCT	CACCTTCCAC	CTGGTGCGGT	GAGCGCGCCC	960
GCTGAACCTC	CCGCTGCTGC	TGCTGCTGCT	GGGGGCCACT	GTGGCCGCCG	AACTCATCTC	1020

بالرامرية

#### PF-0459 US

# (2) INFORMATION FOR SEQ ID NO: 126:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1093 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: PROSNOT16
- (B) CLONE: 2232884
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 126:

AGAGCCCCAG	CCACGCCGGC	CCAGGTGGCC	TCAGGTGAGG	GGGGGCGGAC	GCACCTGTGG	60
GGACGGGACG	ACGAGTTCAA	GCCTCCGTGG	GTGCAGTTGG	TCGCCAGCGA	GGGATGCGGA	120
GACGCCCCTG	AACGACCATG	GCATCGGCCG	ACGAGCTGAC	CTTCCATGAA	TTCGAGGAGG	180
CCACTAATCT	TCTGGCTGAC	ACCCCAGATG	CAGCCACCAC	CAGCAGAAGC	GATCAGCTGA	240
CCCCACAAGG	GCACGTGGCT	GTGGCCGTGG	GCTCAGGTGG	CAGCTATGGA	GCCGAGGATG	300
AGGTGGAGGA	GGAGAGTGAC	AAGGCCGCGC	TCCTGCAGGA	GCAGCAGCAG	CAGCAGCAGC	360
CGGGATTCTG	GACCTTCAGC	TACTATCAGA	GCTTCTTTGA	CGTGGACACC	TCACAGGTCC	420
TGGACCGGAT	CAAAGGCTCA	CTGCTGCCCC	GGCCTGGCCA	CAACTTTGTG	CGGCACCATC	480
TGCGGAATCG	GCCGGATCTG	TATGGCCCCT	TCTGGATCTG	TGCCACGTTG	GCCTTTGTCC	540
TGGCCGTCAC	TGGCAACCTG	ACGCTGGTGC	TGGCCCAGAG	GAGGGACCCC	TCCATCCACT	600
ACAGCCCCCA	GTTCCACAAG	GTGACCGTGG	CAGGCATCAG	CATCTACTGC	TATGCGTGGC	660
TGGTGCCCCT	GGCCCTGTGG	GGCTTCCTGC	GGTGGCGCAA	GGGTGTCCAG	GAGCGCATGG	720
GGCCCTACAC	CTTCCTGGAG	ACTGTGTGCA	TCTACGGCTA	CTCCCTCTTT	GTCTTCATCC	780
CCATGGTGGT	CCTGTGGCTC	ATCCCTGTGC	CTTGGCTGCA	GTGGCTCTTT	GGGGCGCTGG	840
CCCTGGGCCT	GTCAGCCGCC	GGGCTGGTAT	TCACCCTCTG	GCCCGTGGTC	CGTGAGGACA	900
CCAGGCTGGT	GGCCACAGTG	CTGCTGTCCG	TGGTCGTGCT	GCTCCACGCC	CTCCTGGCCA	960
TGGGCTGTAA	GTTGTACTTC	TTCCAGTCGC	TGCCTCCGGA	GAACGTGGCT	CCTCCACCCC	1020
AAATCACATC	TCTGCCCTCA	AACATCGCGC	TGTCCCCTAC	CTTGCCGCAG	TCCCTGGCCC	1080
CCTCCTAGGA	AGG					1093

# (2) INFORMATION FOR SEQ ID NO: 127:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1121 base pairs
    (B) TYPE: nucleic acid
    (C) STRANDEDNESS: single
    (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: COLNNOT11
  - (B) CLONE: 2328134
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 127:

GCGGGGGATG	ACGCCACGGA	CATGGTGGCC	GAGACCGGCG	GGGTGGGGGA	CGTGTCGCGC	60
GGCCGGGTGG	CCTCGGTCGG	TACCCTGGGC	GCGGACAGCT	GCCTCATTAG	TATTCGTACC	120
CACGAGGCGG	CGCAGCGGGC	CCTCGGGGAC	AGCGAGCGTC	GCGGCTATGG	CTTATCACTC	180
GGGCTACGGA	GCCCACGGCT	CCAAGCACAG	GGCCCGGGCA	GCCCCGGATC	CCCCTCCCCT	240
CTTCGATGAC	ACAAGCGGTG	GTTATTCCAG	CCAGCCCGGG	GGATACCCAG	CCACAGGAGC	300
AGACGTGGCC	TTCAGTGTCA	ACCACTTGCT	TGGGGACCCA	ATGGCCAATG	TGGCTATGGC	360
CTATGGCAGC	TCCATCGCAT	CCCATGGGAA	GGACATGGTG	CACAAGGAGC	TGCACCGTTT	420
TGTGTCTGTG	AGCAAACTCA	AGTATTTTT	TGCTGTGGAC	ACAGCCTACG	TGGCCAAGAA	480
GCTAGGGCTG	CTGGTCTTCC	CCTACACACA	CCAGAACTGG	GAAGTGCAGT	ACAGTCGTGA	540
TGCTCCTCTG	CCCCCCGGC	AAGACCTCAA	CGCCCTGAC	CTCTATATCC	CCACGATGGC	600
CTTCATTACT	TACGTGCTCC	TGGCTGGGAT	GGCACTGGGC	ATTCAGAAAA	GGTTCTCCCC	660
GGAGGTGCTG	GGCCTGTGTG	CAAGCACAGC	GCTGGTGTGG	GTGGTGATGG	AGGTGCTGGC	720
CCTGCTCCTG	GGCCTCTACC	TGGCCACCGT	GCGCAGTGAC	CTGAGCACCT	TTCACCTGCT	780
GGCCTACAGT	GGCTACAAAT	ACGTGGGAAT	GATCCTCAGT	GTGCTCACGG	GGCTGCTGTT	840
CGGCAGCGAT	GGCTACTACG	TGGCGCTGGC	CTGGACCTCA	TCGGCGCTCA	TGTACTTCAT	900
TGTGCGCTCT	TTGCGGACAG	CAGCCCTGGG	CCCCGACAGC	ATGGGGGGCC	CCGTCCCCCG	960
GCAGCGTCTC	CAGCTCTACC	TGACTCTGGG	AGCTGCAGCC	TTCCAGCCCC	TCATCATATA	1020
CTGGCTGACT	TTCCACCTGG	TCCGGTGACC	CCCTGGCCCC	AGATGGCACT	GAGTTTTTCA	1080
TTCATTGAAG	ATTTGATTTC	CTTGAAAAAA	AAAAAAAAAG	G		1121

(2) INFORMATION FOR SEQ ID NO: 128:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1861 base pairs

  - (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: ISLTNOT01 (B) CLONE: 2382718
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 128:

(	CGCGGACTGT	GTCTGTTCCC	AGGAGTCCTT	CGGCGGCTGT	TGTGTCAGTG	GCCTGATCGC	60
,	GATGGGGACA	AAGGCGCAAG	TCGAGAGGAA	ACTGTTGTGC	CTCTTCATAT	TGGCGATCCT	120
	GTTGTGCTCC	CTGGCATTGG	GCAGTGTTAC	AGTGCACTCT	TCTGAACCTG	AAGTCAGAAT	180
	TCCTGAGAAT	AATCCTGTGA	AGTTGTCCTG	TGCCTACTCG	GGCTTTTCTT	CTCCCCGTGT	240
	GGAGTGGAAG	TTTGACCAAG	GAGACACCAC	CAGACTCGTT	TGCTATAATA	ACAAGATCAC	300
	AGCTTCCTAT	GAGGACCGGG	TGACCTTCTT	GCCAACTGGT	ATCACCTTCA	AGTCCGTGAC	360
	ACGGGAAGAC	ACTGGGACAT	ACACTTGTAT	GGTCTCTGAG	GAAGGCGGCA	ACAGCTATGG	420
	GGAGGTCAAG	GTCAAGCTCA	TCGTGCTTGT	GCCTCCATCC	AAGCCTACAG	TTAACATCCC	480
	CTCCTCTGCC	ACCATTGGGA	ACCGGGCAGT	GCTGACATGC	TCAGAACAAG	ATGGTTCCCC	540
	ACCTTCTGAA	TACACCTGGT	TCAAAGATGG	GATAGTGATG	CCTACGAATC	CCAAAAGCAC	600
	CCGTGCCTTC	AGCAACTCTT	CCTATGTCCT	GAATCCCACA	ACAGGAGAGC	TGGTCTTTGA	660
	TCCCCTGTCA	GCCTCTGATA	CTGGAGAATA	CAGCTGTGAG	GCACGGAATG	GGTATGGGAC	720
	ACCCATGACT	TCAAATGCTG	TGCGCATGGA	AGCTGTGGAG	CGGAATGTGG	GGGTCATCGT	780
	GGCAGCCGTC	CTTGTAACCC	TGATTCTCCT	GGGAATCTTG	GTTTTTGGCA	TCTGGTTTGC	840
	CTATAGCCGA	GGCCACTTTG	ACAGAACAAA	GAAAGGGACT	TCGAGTAAGA	AGGTGATTTA	900
	CAGCCAGCCT	AGTGCCCGAA	. GTGAAGGAGA	ATTCAAACAG	ACCTCGTCAT	TCCTGGTGTG	960
	AGCCTGGTCG	GCTCACCGCC	TATCATCTGC	ATTTGCCTTA	CTCAGGTGCT	ACCGGACTCT	1020
	GGCCCCTGAT	GTCTGTAGTT	' TCACAGGATG	CCTTATTTGT	CTTCTACACC	CCACAGGGCC	1080
	CCCTACTTCT	TCGGATGTGT	TTTTAATAAT	GTCAGCTATG	TGCCCCATCC	TCCTTCATGC	1140
	CCTCCCTCCC	TTTCCTACCA	CTGCTGAGTG	GCCTGGAACT	TGTTTAAAGT	GTTTATTCCC	1200
	CATTTCTTTG	AGGGATCAGG	AAGGAATCC1	GGGTATGCCA	TTGACTTCCC	TTCTAAGTAG	1260
	ACAGCAAAAA	TGGCGGGGG	CGCAGGAATO	TGCACTCAAC	TGCCCACCTG	GCTGGCAGGG	1320
	ATCTTTGAAT	AGGTATCTTC	AGCTTGGTT	TGGGCTCTTI	CCTTGTGTAC	TGACGACCAG	1380
	GGCCAGCTGT	TCTAGAGCG	GAATTAGAGO	CTAGAGCGGC	TGAAATGGTI	GTTTGGTGAT	1440
	GACACTGGGG	TCCTTCCATO	C TCTGGGGCC	C ACTCTCTTCT	GTCTTCCCAT	GGGAAGTGCC	: 1500

# ACTGGGATCC CTCTGCCTG TCCTCTGAA TACAAGCTGA CTGACATTGA CTGTGTCTGT 1560 GGAAAATGGG AGCTCTTGTT GTGGAGAGCA TAGTAAATTT TCAGAGAACT TGAAGCCAAA 1620 AGGATTTAAA ACCGCTGCTC TAAAGAAAAG AAAACTGGAG GCTGGGCGCA GTGGCTCACG 1680 CCTATAATCC CAGAGGCTGA GGCAGGCGGA TCACCTGAGG TCGGGAGTTC GGGATCAGCC 1740

CTGTAATCCC AGCTGCTCAG GAGCCTGGCA ACAAGAGCAA AACTCCAGCT CAAAAAAAAA 1860

TGACCAACAT GGAGAAACCC TACTGAGAAT ACAAAGTTAG CCAGGCATGG TGGTGCATGC 1800

A 1861

# (2) INFORMATION FOR SEQ ID NO: 129:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1975 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: ENDANOT01
  - (B) CLONE: 2452208
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 129:

GTTTGGAGGA	GACTCGGATA	TACCTTCTCA	GAAGCTGCAC	AGGAGGAAAG	CAGTGACAAA	60
GAAAGAAGTT	GTCATTCTTT	GCACGAAACT	GGATGGCTTC	TACAGGGAGC	CAGGCCTCTG	120
ATATAGACGA	GATTTTTGGA	TTCTTCAACG	ATGGCGAACC	TCCCACCAAA	AAGCCCAGGA	180
AGCTGCTTCC	AAGCTTAAAA	ACTAAGAAGC	CTCGAGAACT	TGTGCTAGTG	ATTGGAACAG	240
GCATTAGTGC	TGCAGTTGCG	CCCCAAGTTC	CAGCCCTCAA	ATCCTGGAAG	GGGTTAATTC	300
AGGCCTTACT	GGATGCTGCC	ATTGATTTTG	ATCTTTTAGA	AGATGAGGAG	AGCAAAAAGT	360
TTCAGAAATG	TCTCCATGAA	GACAAGAACC	TGGTCCATGT	TGCCCATGAC	CTTATCCAGA	420
AACTCTCTCC	TCGTACCAGT	AATGTTCGAT	CCACATTTTT	CAAGGACTGT	TTATATGAAG	480
TATTTGATGA	CTTGGAGTCA	AAGATGGAAG	ATTCTGGAAA	ACAGCTACTT	CAGTCAGTTC	540
TCCACCTGAT	GGAAAATGGA	GCCCTCGTAT	TAACTACAAA	TTTTGATAAT	CTCTTGGAAC	600
TGTATGCAGC	AGATCAGGGG	AAACAGCTTG	AATCCCTTGA	CCTTACTGAT	GAGAAAAAGG	660
TCCTCGAGTG	GGCTCAGGAG	AAGCGTAAGC	TGAGCGTGTT	GCATATTCAC	GGAGTCTACA	720
CCAACCCTAG	TGGCATTGTC	CTTCATCCGG	CTGGATATCA	GAACGTGCTC	AGGAACACTG	780
AAGTCATGAG	AGAAATTCAG	AAACTCTACG	AAAACAAGTC	ATTTCTTTTC	CTGGGCTGTG	840
GCTGGACTGT	GGATGACACC	ACTTTCCAGG	CCCTTTTCTT	GGAGGCTGTC	AAGCATAAAT	900

CTGACCTAGA ACATTICATG CTGGTTCGGA GAGGAGACGT AGATGAGTTC AAAAAGCTTC 960 GAGAAAACAT GCTGGACAAG GGGATTAAAG TCATCTCCTA TGGAGATGAC TATGCCGATC 1020 TTCCAGAATA TTTCAAGCGA CTGACATGTG AGATCTCCAC AAGGGGTACA TCAGCAGGGA 1080 TGGTGAGAGA AGGTCAGCTA AATGGCTCAT CTGCAGCACA CAGTGAAATA AGAGGCTGTA 1140 GTACATGAGC GAGCTAGAGA AATCACCACC GTTTAGACCA AGCTGTAAGG CCCTACTACA 1200 GACAGTGTTT AACAAGTAAA CTTACAAGAA CCCAACACAA TTCCCAGAAA GTAACAATAG 1260 CCAGAGGTTG AAGGGCGGGG TAGAAGAGGG GGGAATGTTG CAGCGTAATC CTTCATACCA 1320 CCTGGTTCTT GATATTCTGC CGCCTGTTCA AGTTCAAGAA TAAAAGCGAC AGCAGGACCC 1380 AAATGCAGCT CCCAACCCAC TCCCCAGGCT AGACATGCTT GTGTCCACAC AGCACACCAA 1440 TGTGATACTT CCACTGACCG GCTGCAGCTC TGCATGAAGG ACTCGGGGTC TGGATGCCAT 1500 GGAATCACTG TGGCTCTTGT TGCAGTTTTG TACTCTATAC TTGGTTTTTC AATTAAGCTT 1560 AATGGCTTTT TTAAAACATG ACTTGAAGCT CTAGTTTTCT AGATCTTTTA CAGTGTACAG 1620 TATTTTACAT AACTAAGCTG TATTAAAAGC TTGTTCATTT ACTTGCCAGG ACCCTGGCTC 1680 TACTTTTAGA GTCATTGTAA GAAACTCTAA CTTGCATCAA GGTACTAATA AGCTTAATTT 1740 TAATAACCCA AAGTTTAAAG GTTCCGATCT TTCTCCTTGG GGTGGAGTGA TCTCATTCTC 1800 AGGACAACCG TTTACTTACC TGATTCCTCG GAGCATTATC AACTTCTGCT CTGTTGTCCT 1860 GACCATACAT ATGTCCTAGA ACTACAGTTA AGTGTGTTGT GGAATTTTAG TTTTGAATCC 1920 GGAATAAATG AAGTCCCAGG ACTCAAAGAA GAGAGAAAAA AAAAAAGGGG GCCCC

# (2) INFORMATION FOR SEQ ID NO: 130:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2160 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: ENDANOT01
  - (B) CLONE: 2457825
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 130:

TCTACTGTCC CCTGCCCTGT ACCCCCAGGC ATTGATCTGG AGAACATTGT GTACTACAAG 60

GACGACACCC ACTACTTTGT GATGACAGCC AAGAAGCAGT GCCTGCTGCG GCTGGGGGTG 120

CTGCGCCAGG ACTGGCCAGA CACCAATCGG CTGCTGGGCA GTGCCAATGT GGTGCCCGAG 180

GCTCTGCAGC GCTTTACCCG GGCAGCTGCT GACTTTGCCA CCCATGGCAA GCTCGGGAAA 240

CTAGAGTTTG	CCCAGGATGC	CCATGGGCAG	CCTGATGTCT	CTGCCTTTGA	CTTCACGAGC	300
ATGATGCGGG	CAGAGAGTTC	TGCTCGTGTG	CAAGAGAAGC	ATGGCGCCCG	CCTGCTGCTG	360
GGACTGGTGG	GGGACTGCCT	GGTGGAGCCC	TTCTGGCCCC	TGGGCACTGG	AGTGGCACGG	420
GGCTTCCTGG	CAGCCTTTGA	TGCAGCCTGG	ATGGTGAAGC	GGTGGGCAGA	GGGCGCTGAG	480
TCCCTAGAGG	TGTTGGCTGA	GCGTGAGAGC	CTGTACCAGC	TTCTGTCACA	GACATCCCCA	540
GAAAACATGC	ATCGCAATGT	GGCCCAGTAT	GGGCTGGACC	CAGCCACCCG	CTACCCCAAC	600
CTGAACCTCC	GGGCAGTGAC	CCCCAATCAG	GTACGAGACC	TGTATGATGT	GCTAGCCAAG	660
GAGCCTGTGC	AGAGGGACAA	CGACAAGACA	GATACAGGGA	TGCCAGCCAC	CGGGTCGGCA	720
GGCACCCAGG	AGGAGCTGCT	ACGCTGGTGC	CAGGAGCAGA	CAGCTGGGTA	CCCGGGAGTC	780
CACGTCTCCG	ATTTGTCTTC	CTCCTGGGCT	GATGGGCTAG	CTCTGTGTGC	CCTGGTGTAC	840
CGGCTGCAGC	CTGGCCTGCT	GGAACCCTCA	GAGCTGCAGG	GGCTGGGAGC	TCTGGAAGCA	900
ACTGCTTGGG	CACTAAAGGT	GGCAGAGAAT	GAGCTGGGCA	TCACACCGGT	GGTGTCTGCA	960
CAGGCCGTGG	TAGCAGGGAG	TGACCCACTG	GGCCTCATTG	CCTACCTCAG	CCACTTCCAC	1020
AGTGCCTTCA	AGAGCATGGC	CCACAGCCCA	GGCCCTGTCA	GCCAGGCCTC	CCCAGGGACC	1080
TCCAGTGCTG	TATTATTCCT	TAGTAAACTT	CAGAGGACCC	TGCAGCGATC	CCGGGCCAAG	1140
GAAAATGCAG	AGGATGCTGG	TGGCAAGAAG	CTGCGCTTGG	AGATGGAGGC	CGAGACCCCA	1200
AGTACTGAGG	TGCCACCTGA	CCCAGAGCCT	GGTGTACCCC	TGACACCCC	ATCCCAACAC	1260
CAGGAGGCCG	GTGCTGGGGA	CCTGTGTGCA	CTTTGTGGGG	AACACCTCTA	TGTCCTGGAA	1320
CGCCTCTGTG	TCAACGGCCA	TTTCTTCCAC	CGGAGCTGCT	TCCGCTGCCA	TACCTGTGAG	1380
GCCACACTGT	GGCCAGGTGG	CTACGAGCAG	CACCCAGGCA	GTAGAACGTC	TCAGTTCTTC	1440
TTCTCAGCTC	TTGTGGCCAT	GGAGAAGGAG	GAAAAAGAGA	GTCCCTTCTC	CAGTGAAGAG	1500
GAAGAAGAAG	ATGTGCCTTT	GGACTCAGAT	GTGGAACAGG	CCCTGCAGAC	CTTTGCCAAG	1560
ACCTCAGGCA	CCATGAATAA	CTACCCAACA	TGGCGTCGGA	CTCTGCTGCG	CCGTGCGAAG	1620
GAGGAGGAGA	. TGAAGAGGTT	CTGCAAGGCC	CAGACCATCC	AACGGCGACT	AAATGAGATT	1680
GAGGCTGCCT	TGAGGGAGCT	AGAGGCCGAG	GGCGTGAAGC	TGGAGCTGGC	CTTGAGGCGC	1740
CAGAGCAGTT	CCCCAGAACA	. GCAAAAGAAA	CTATGGGTAG	GACAGCTGCT	' ACAGCTCGTT	1800
GACAAGAAAA	ACAGCCTGGT	GGCTGAGGAG	GCCGAGCTCA	TGATCACGGT	GCAGGAATTG	1860
AATCTGGAGG	AGAAACAGTG	GCAGCTGGAC	CAGGAGCTAC	GAGGCTACAT	GAACCGGGAA	1920
GAAAACCTAA	AGACAGCTGC	: TGATCGGCAG	GCTGAGGACC	AGGTCCTGAG	GAAGCTGGTG	1980
GATTTGGTCA	ACCAGAGAGA	TGCCCTCATC	CGCTTCCAGG	aggagcgca	GCTCAGCGAG	2040
CTGGCCTTGG	GGACAGGGG	CCAGGGCTAG	ACGAGGGTGG	GCCGTCTGC	TTCGTTCCCA	2100

CAAAGAAAGC ACCTCACCCC AGCACAGTGC CACCCCTGTT CATCTGGGCT GCCTGGCAGA 2160

- (2) INFORMATION FOR SEQ ID NO: 131:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 546 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: THP1NOT03
    - (B) CLONE: 2470740
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 131:
- (2) INFORMATION FOR SEQ ID NO: 132:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 581 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: SMCANOT01
    - (B) CLONE: 2479092
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 132:
- GCCATGGAGG CCCTGAGGAG GGCCCACGAG GTCGCGCTCC GCCTGCTGCT GTGTAGGCCG 60
  TGGGCCTCGC GCGCCGCGC CCGCCCCAAG CCCAGCGCCT CGGAGGTGCT GACGCGCAT 120

# CTGCTGCAGC GGCGCTGCC GCACTGGACC TCCTTCTGCG TGCCCTACAG CGCCGTCCGC 180 AACGACCAGT TCGGCCTCTC GCACTTCAAC TGGCCGGTGC AGGGCGCCAA CTACCACGTC 240 CTGCGCACCG GCTGCTTCCC CTTCATCAAG TACCACTGCT CCAAGGCTCC CTGGCAGGAC 300 CTGGCCCGGC AGAACCGCTT CTCACGGCG CTCAAGGTCG TCAACCTCGG TATTCCAACT 360 TTATTATATA GACTTGGCTC CTGGTTATTT GCCAGAGTCA CAGAGACTGT GCATACCAGT 420 TATGGACCCA TAACAGTTTA TTTTCTCAAT AAAGAAGATG AAGGTGCCAT GTATTGAAAG 480 TGTGCGTCAA AGAACATAAA TATCAGTGGA TTTTCTCTGT GTATATGTGC AGTATTTATT 540

581

# (2) INFORMATION FOR SEQ ID NO: 133:

PF-0459 US

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1259 base pairs

TTTGATCCTT TAAAATAAAA CTTTTGCAAA TAAAAAAAAA A

- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: SMCANOT01
  - (B) CLONE: 2480544
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 133:

GGGCTGGGCC	CCGCCGCAGC	TCCAGCTGGC	CGGCTTGGTC	CTGCGGTCCC	TTCTCTGGGA	60
GGCCCGACCC	CGGCCGCGCC	CAGCCCCCAC	CATGCCACCC	GCGGGGCTCC	GCCGGGCCGC	120
GCCGCTCACC	GCAATCGCTC	TGTTGGTGCT	GGGGGCTCCC	CTGGTGCTGG	CCGGCGAGGA	180
CTGCCTGTGG	TACCTGGACC	GGAATGGCTC	CTGGCATCCG	GGGTTTAACT	GCGAGTTCTT	240
CACCTTCTGC	TGCGGGACCT	GCTACCATCG	GTACTGCTGC	AGGGACCTGA	CCTTGCTTAT	300
CACCGAGAGG	CAGCAGAAGC	ACTGCCTGGC	CTTCAGCCCC	AAGACCATAG	CAGGCATCGC	360
CTCAGCTGTG	ATCCTCTTTG	TTGCTGTGGT	TGCCACCACC	ATCTGCTGCT	TCCTCTGTTC	420
CTGTTGCTAC	CTGTACCGCC	GGCGCCAGCA	GCTCCAGAGC	CCATTTGAAG	GCCAGGAGAT	480
TCCAATGACA	GGCATCCCAG	TGCAGCCAGT	ATACCCATAC	CCCCAGGACC	CCAAAGCTGG	540
CCCTGCACCC	CCACAGCCTG	GCTTCATGTA	CCCACCTAGT	GGTCCTGCTC	CCCAATATCC	600
ACTCTACCCA	GCTGGGCCCC	CAGTCTACAA	CCCTGCAGCT	CCTCCTCCCT	ATATGCCACC	660
ACAGCCCTCT	TACCCGGGAG	CCTGAGGAAC	CAGCCATGTC	TCTGCTGCCC	CTTCAGTGAT	720
GCCAACCTTG	GGAGATGCCC	TCATCCTGTA	CCTGCATCTG	GTCCTGGGGG	TGGCAGGAGT	780
CCTCCAGCCA	CCAGGCCCCA	GACCAAGCCA	AGCCCTGGGC	CCTACTGGGG	ACAGAGCCCC	840

\$ · ;

AGGGAAGTGG AACAGGAGCT GAACTAGAAC TATGAGGGGT TGGGGGAGG GCTTGGAATT 900
ATGGGCTATT TTTACTGGGG GCAAGGGAGG GAGATGACAG CCTGGGTCAC AGTGCCTGTT 960
TTCAAATAGT CCCTCTGCTC CCAAGATCCC AGCCAGGAAG GCTGGGGCCC TACTGTTTGT 1020
CCCCTCTGGG CTGGGGTGGG GGGAGGGAGG AGGTTCCGTC AGCAGCTGGC AGTAGCCCTC 1080
CTCTCTGGCT GCCCCACTGG CCACATCTCT GGCCTGCTAG ATTAAAGCTG TAAAGACATA 1140
ACTCATATCA GTCGCATCAT TGGACCCATC CACACCTTCC AGGAACACCG NCTTCAGCTG 1200
GGCCCAGACT GTTGCCCACT CCATATTCCA AAAGTAGGGG AGGGCCAGCA CCAGCATCG 1259

# (2) INFORMATION FOR SEQ ID NO: 134:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2033 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRAITUT21
- (B) CLONE: 2518547

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 134:

CGGCTCGAGG	CCGCAGCCCC	ATGGACAGTC	TTCTGCACCC	CCGGGAGCGC	CCTGGATCCA	60
CTGCCTCCGA	GAGCTCAGCC	TCTCTGGGCA	GTGAGTGGGA	CCTCTCAGAA	TCTTCTCTCA	120
GCAACCTGAG	TCTTCGCCGT	TCCTCAGAGC	GCCTCAGTGA	CACCCCTGGA	TCCTTCCAGT	180
CACCTTCCCT	GGAAATTCTG	CTGTCCAGCT	GCTCCCTGTG	CCGTGCCTGT	GATTCGCTGG	240
TGTATGATGA	GGAAATCATG	GCTGGCTGGG	CACCTGATGA	CTCTAACCTC	AACACAACCT	300
GCCCCTTCTG	CGCCTGCCCC	TTTGTGCCCC	TGCTCAGTGT	CCAGACCCTT	GATTCCCGGC	360
CCAGTGTCCC	CAGCCCCAAA	TCTGCTGGTG	CCAGTGGCAG	CAAAGATGCT	CCTGTCCCTG	420
GTGGTCCTGG	CCCTGTGCTC	AGTGACCGAA	GGCTCTGCCT	TGCTCTGGAT	GAGCCCAGCT	480
CTGCAACGGG	CACATGGGGG	GAGCCTCCCG	GCGGGTTGAG	AGTGGGGCAT	GGGCATACCT	540
GAGCCCCCTG	GTGCTGCGTA	AGGAGCTGGA	GTCGCTGGTA	GAGAACGAGG	GCAGTGAGGT	600
GCTGGCGTTG	CCTGAACTGC	CCTCTGCCCA	CCCCATCATC	TTCTGGAACC	TTTTGTGGTA	660
TTTCCAACGG	CTACGCCTGC	CCAGTATTCT	ACCAGGCCTG	GTGCTGGCCT	CCTGTGATGG	720
GCCTTCGCAC	TCCCAGGCCC	CATCTCCTTG	GCTAACCCCT	GATCCAGCCT	CTGTTCAGGT	780
ACGGCTGCTG	TGGGATGTAC	TGACCCCTGA	CCCCAATAGC	TGCCCACCTC	TCTATGTGCT	840
CTGGAGGGTC	CACAGCCAGA	TCCCCCAGCG	GGTGGTATGG	CCAGGCCCTG	TACCTGCATC	900

CCTTAGTTTG GCACTGTTGG AGTCAGTGCT GCGCCATGTT GGACTCAATG AAGTGCACAA 960 GGCTGTGGGG CTCCTGCTGG AAACTCTAGG GCCCCCACCC ACTGGCCTGC ACCTGCAGAG 1020 GGGAATCTAC CGTGAGATAT TATTCCTGAC AATGGCTGCT CTGGGCAAGG ACCACGTGGA 1080 CATAGTGGCC TTCGATAAGA AGTACAAGTC TGCCTTTAAC AAGCTGGCCA GCAGCATGGG 1140 CAAGGAGGAG CTGAGGCACC GGCGGGCGCA GATGCCCACT CCCAAGGCCA TTGACTGCCG 1200 AAAATGTTTT GGAGCACCTC CAGAATGCTA GAGACCTTAA GCTTCCCTCT CCAGCCTAGG 1260 GTGGGGAAGT GAGGAAGAAG GGATTCTAGA GTTAAACTGC CTCCCTGTTG CCTTCATGGA 1320 GTTGGGAACA GGCTGGGAAG GATGCCCAGT CAAAGGCTCC AAGCGAGGAC AACAGGAAGA 1380 GGGATCCACT GTTACCAAAA GTCCTGATTC CCCCATCACC AACCTACCCA GTTTGTTCGT 1440 GCTGATGTTG GGGGGAGTCT GGGGGGGAGTT GGTACAGCTC TGTTCTTCCC TTGTCCTATA 1500 CCGGGAACTC CCCTCCAGGG TACCCACAGA TCTGCATTGC CCTGGTCATT TTAGAAGTTT 1560 TTGTTTTAAA AAACAACTGG AAAGATGCAG AGCTACTGAG CCTTTGCCCT GAATGGGAGG 1620 TAGGGATGTC ATTCTCCACC AATAATGGTC CCTCTTCCCT GACGTTGCTG AAGGAGCCCA 1680 AGGCTCTCCA TGCCTTTCTA CCTAAGTGTT TGTATTTTAT TTTAAATTAT TTATTCTGGA 1740 GCCACAGCCC CCTTGCTTAT GAGGTTCTTA TGGAGAGTGA GAAAGGGAAG GGAAATAGGG 1800 CACCATGGTC CGGTGGTTTG TAGTTCCTTC AAAGTCAGGC ACTGGGAGCT AGAGGAGTCT 1860 CAAGCTCCCC TTAGGAAGAA CTGGTGCCCC CTCCAGTCCT AATTTTTCTT GCCTGCCCCG 1920 CCTTGGGGAA TGCCTCACCC ACCCAGGTCC TGACCTGTGC AATAAGGATT GTTCCCTGCG 1980 AAGTTTTGTT GGATGTAAAT ATAGTAAAAG CTGCTTCTGT CTTTTTCAAA AAA 2033

# (2) INFORMATION FOR SEQ ID NO: 135:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 3007 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: GBLANOT02
  - (B) CLONE: 2530650
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 135:

GCCCACTGGG CTCTCCCGGC TGCAGTGCCA GGGCGCAGGA CGCGGCCGAT CTCCCGCTCC 60

CGCCACCTCC GCCACCATGC TGCTCCCCCA GCTCTGCTGG CTGCCGCTGC TCGCTGGGCT 120

GCTCCCGCCG GTGCCCGCTC AGAAGTTCTC GGCGCTCACG TTTTTGAGAG TGGATCAAGA 180

TAAAGACAAG	GATTGTAGCT	TGGACTGTGC	GGGTTCGCCC	CAGAAACCTC	TCTGCGCATC	240
TGACGGAAGG	ACCTTCCTTT	CCCGTTGTGA	ATTTCAACGT	GCCAAGTGCA	AAGATCCCCA	300
GCTAGAGATT	GCATATCGAG	GAAACTGCAA	AGACGTGTCC	AGGTGTGTGG	CCGAAAGGAA	360
GTATACCCAG	GAGCAAGCCC	GGAAGGAGTT	TCAGCAAGTG	TTCATTCCTG	AGTGCAATGA	420
CGACGGCACC	TACAGTCAGG	TCCAGTGTCA	CAGCTACACG	GGATACTGCT	GGTGCGTCAC	480
GCCCAACGGG	AGGCCCATCA	GCGGCACTGC	CGTGGCCCAC	AAGACGCCCC	GGTGCCCGGG	540
TTCCGTAAAT	GAAAAGTTAC	CCCAACGCGA	AGGCACAGGA	AAAACAGATG	ATGCCGCAGC	600
TCCAGCGTTG	GAGACTCAGC	CTCAAGGAGA	TGAAGAAGAT	ATTGCATCAC	GTTACCCTAC	660
CCTTTGGACT	GAACAGGTTA	AAAGTCGGCA	GAACAAAACC	AATAAGAATT	CAGTGTCATC	720
CTGTGACCAA	GAGCACCAGT	CTGCCCTGGA	GGAAGCCAAG	CAGCCCAAGA	ACGACAATGT	780
GGTGATCCCT	GAGTGTGCGC	ACGGCGGCCT	CTACAAGCCA	GTGCAGTGCC	ACCCCTCCAC	840
GGGGTACTGC	TGGTGCGTCC	TGGTGGACAC	GGGGCGCCCC	ATTCCCGGCA	CATCCACAAG	900
GTACGAGCAG	CCGAAATGTG	ACAACACGGG	CCAGGGCCCA	CCCAGCCAAA	GCCCGGGACC	960
TGTACAAGGG	CCGCCAGCTA	CAAGGTTGTC	CGGGTGCCAA	AAAGCATGAG	TTTCTGACCA	1020
GCGTTCTGGA	CGCGCTGTCC	ACGGACATGG	TCCACGCCGC	CTCCGACCCC	TCCTCCTCGT	1080
CAGGCAGGCT	CTCAGAACCC	GACCCCAGCC	ATACCCTAGA	GGAGCGGGTG	GTGCACTGGT	1140
ACTTCAAACT	ACTGGATAAA	AACTCCAGTG	GAGACATCGG	CAAAAAGGAA	ATCAAACCCT	1200
TCAAGAGGTT	CCTTCGCAAA	AAATCAAAGC	CCAAAAAATG	TGTGAAGAAG	TTTGTTGAAT	1260
ACTGTGACGT	GAATAATGAC	AAATCCATCT	CCGTACAAGA	ACTGATGGGC	TGCCTGGGCG	1320
TGGCGAAAGA	GGACGGCAAA	GCGGACACCA	AGAAACGCCA	CACCCCCAGA	GGTCATGCTG	1380
AAAGTACGTC	TAATAGACAG	CCAAGGAAAC	AAGGATAAAT	GGCTCATACC	CCGAAGGCAG	1440
TTCCTAGACA	. CATGGGAAAT	TTCCCTCACC	C AAAGAGCAAT	TAAGAAAAC <i>P</i>	AAAACAGAAA	1500
CACATAGTAI	TTGCACTTTG	TACTTTAAAT	GTAAATTCAC	TTTGTAGAAA	TGAGCTATTT	1560
AAACAGACTO	TTTTAATCTG	TGAAAATGGA	A GAGCTGGCTT	CAGAAAATTA	A ATCACATACA	1620
ATGTATGTGT	CCTCTTTTGA	CCTTGGAAAT	CTGTATGTG	G TGGAGAAGTA	A TTTGAATGCA	1680
TTTAGGCTTA	ATTTCTTCGC	CTTCCACATO	TTAACAGTAG	G AGCTCTATGO	C ACTCCGGCTG	1740
CAATCGTATC	G GCTTTCTCTA	ACCCCTGCAG	G TCACTTCCAC	G ATGCCTGTG	C TTACAGCATI	1800
GTGGAATCAT	GTTGGAAGCT	CCACATGTCC	C ATGGAAGTT	r grgargraco	G GCCGACCCTA	1860
CAGGCAGTT	A ACATGCATGG	GCTGGTTTG	T TTCTTGGGA	TTTCTGTTA	TTTGTCTTGT	1920
TTTGCTTTC	C AGAGATCTTG	CTCATACAA	r gaatcacgc	A ACCACTAAA	G CTATCCAGTT	1980
AAGTGCAGG:	r AGTTCCCCTG	G GAGGAAATA	A TATTTTCAA	A CTGTCGTTG	G TGTGATACTI	2040

TGGCTCAAAG GATCTTTGCT TTTCCATTTT AAGCTTCTGT TTTGAGTTTT GCCCTGGGGC 2100 TTGAATGAGT CCCAGAGAGT CGTTCGGATG GTGGGAGGCT GCCTAGGAGG CAGTAAATCC 2160 AGTCACAGTG CCTGGGAGGG GCCCATCCTT CCAAAATGTA AATCCAGTCG CGGTGTGACC 2220 GAGCTGGCTA ACAGGCTTGT CTGCCTGGTT TTCCTCCTAC ACGTGGACAT TATTCTCCTG 2280 ATCCTCCTAC CTGGTCCACC CCAGGGCTAC CGGAAGGTAA AATCTTCACC TGAACCAATT 2340 ATGAGCAGTC TCCTTACTGA AGGTACAGCC GGATACGTGG TGCCCCCGGG GCTGGTGTTG 2400 GCAGCCGGGG GGAGGTGCCT GAGGGTCCCC ACGGTTCCTT TCTGCTTTTC TGAATGCATC 2460 AAGGGTACGA GAACTTGCCA ATGGGAAATT CATCCGAGTG GCACTGGCAG AGAAGGATAG 2520 GAGTGGAATG CCCACACAGT GACCAACAGA ACTGGTCTGC GTGCATAACC AGCTGCCACC 2580 CTCAGGCCTG GGCCCCAGAG CTCAGGGCAC CCAGTGTCTT AAGGAACCAT TTGGAGGACA 2640 GTCTGAGAGC AGGAACTTCA AGCTGTGATT CTATCTCGGC TCAGACTTTT GGTTGGAAAA 2700 AGATCTTCAT GGCCCCAAAT CCCCTGAGAC ATGCCTTGTA GAATGATTTT GTGATGTTGT 2760 GATGCTTGTG GAGCATCGCG TAAGGCTTCT TGCTTATTTA AACTGTGCAA GGTAAAAATC 2820 AAGCCTTTGG AGCCACAGAA CCAGCTCAAG TACATGCCAA TGTTGTTTAA GAAACAGTTA 2880 TGATCCTAAA CTTTTTGGAT AATCTTTTAT ATTTCTGACC TTTGAATTTA ATCATTGTTC 2940 TTAGATTAAA ATAAAATATG CTATTGAAAC TAAAAAAAAA AAAGAGGGGA GAAGAAAAA 3000 AAAAAGG 3007

# (2) INFORMATION FOR SEQ ID NO: 136

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1229 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: THYMNOT04
  - (B) CLONE: 2652271
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 136:

CTCTCTGCTC CGGTGCAGGC CCGCAGGCGC CCTGGGCTGG GAGCAACGCG ACTGACCGTG 60
GTCGTGGGCG GACGGCGCT GCAGCGTGGA GGAGCTGGGG TCGCTGTGGG TCGCGAACAG 120
AGCCCGGGAC GTGCGCGCTT GGTGCACGAT CCTGAAGGGG AGCTCCGAGG GGCCCGGGTC 180
TCCAGGGCTG CTGCGGCCAT TCCCGGAGCC CGGCGGGGG CCCGCGAGAT ACTGGTTTAG 240
GCCGTCCCAG GGCTCCGGGC GCACCCGGTG GCCGCTGCTG CAGCGGAGGG AGCGCGGCGG 300

CGCGGGGGCT	CGGAGACAGC	GTTTCTCCCG	GAAGTCTTCC	TCGGGCAGCA	GGTGGGAAGT	360
GGGAGCCGGA	GCGGCAGCTG	GCAGCGTTCT	CTCCGCAGGT	CGGCACCATG	CGCCCTGCAG	420
CCCTGCGCGG	GGCCCTGCTG	GGCTGCCTCT	GCCTGGCGTT	GCTTTGCCTG	GGCGGTGCGG	480
ACAAGCGCCT	GCGTGACAAC	CATGAGTGGA	AAAAACTAAT	TATGGTTCAG	CACTGGCCTG	540
AGACAGTATG	CGAGAAAATT	CAAAACGACT	GTAGAGACCC	TCCGGATTAC	TGGACAATAC	600
ATGGACTATG	GCCCGATAAA	AGTGAAGGAT	GTAATAGATC	GTGGCCCTTC	AATTTAGAAG	660
AGATTAAGGA	TCTTTTGCCA	GAAATGAGGG	CATACTGGCC	TGACGTAATT	CACTCGTTTC	720
CCAATCGCAG	CCGCTTCTGG	AAGCATGAGT	GGGAAAAGCA	TGGGACCTGC	GCCGCCCAGG	780
TGGATGCGCT	CAACTCCCAG	AAGAAGTACT	TTGGCAGAAG	CCTGGAACTC	TACAGGGAGC	840
TGGACCTCAA	CAGTGTGCTT	CTAAAATTGG	GGATAAAACC	ATCCATCAAT	TACTACCAAG	900
TTGCAGATTT	TAAAGATGCC	CTTGCCAGAG	TATATGGAGT	GATACCCAAA	ATCCAGTGCC	960
TTCCACCAAG	CCAGGATGAG	GAAGTACAGA	CAATTGGTCA	GATAGAACTG	TGCCTCACTA	1020
AGCAAGACCA	GCAGCTGCAA	AACTGCACCG	AGCCGGGGGA	GCAGCCGTCC	CCCAAGCAGG	1080
AAGTCTGGCT	GGCAAATGGG	GCCGCCGAGA	GCCGGGGTCT	GAGAGTCTGT	GAAGATGGCC	1140
CAGTCTTCTA	TCCCCCACCT	AAAAAGACCA	AGCATTGATG	CCCAAGTTTT	GGAAATATTC	1200
TGTTTTAAAA	AGCATGAGGT	AGGCATGTC				1229

# (2) INFORMATION FOR SEQ ID NO: 137:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1972 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGTUT11
- (B) CLONE: 2746976
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 137:

ACAGGGCTT CCCCTTCGCC GCCGCCCG CCGCCCGA AGCTCCGCCG CGCCCGCCG 60

CCGCGGCCGC CATGCAGTTT ATGTTGCTTT TTAGTCGTCA GGGAAAGCTT CGACTGCAAA 120

AATGGTATGT CCCACTATCA GACAAAGAGA AGAGAAAGAT CACAAGAGAA CTTGTTCAGA 180

CCGTTTTAGC ACGGAAACCT AAAATGTGCA GCTTCCTTGA GTGGCGAGAT CTGAAGATTG 240

TTAATTACCCT GGAAATAATT CATCGTTATG TGGAATTACT TGACAAGTAT TTCGGCAGTG 360

TCTGTGAACT	AGATATCATC	TTTAATTTTG	AGAAGGCTTA	TTTTATTTTG	GATGAGTTTC	420
TTTTGGGAGG	GGAAGTTCAG	GAAACATCCA	AGAAAAATGT	CCTTAAAGCA	ATTGAGCAGG	480
CTGATCTACT	GCAGGAGGAT	GCGAAAGAAG	CTGAAACCCC	ACGTAGTGTT	CTTGAAGAAA	540
TTGGACTGAC	ATAACTCTCC	TCCCTTGTTG	ATGACTTCTT	GTGGCATTTC	ACACACTGTA	600
GATGGTCACT	CCCTTCATGT	CCATGTTAGC	TCATGGTGTA	AGATGATGTC	TTGTCAGTAT	660
TACTGTTTTG	CTAAGCCGCT	TCATTCATGC	CTACACAATT	TTTTTTTAAA	AGGGAACTTT	720
AGTTAATTAA	GTGATAAGGG	ACTTAAATAT	GAATTAGAAT	GGTGCAGAAA	GAGATACCTT	780
TTCTGGATAT	TTTAAAGTTT	AAAGGTCAGT	TTCTCTTAAT	CTGATTATGT	GCACATATGA	840
AAATGGCACA	TCATATACAT	GTAAAATCAG	GCAGTATACA	TTTATTAATT	ACTGTATTTG	900
ACAAAGGAAA	CTCTTAAATT	ATAATGTGAA	ACCTGGTTTT	ATGAAACCAA	AGACTAGTGC	960
AGCATTTCAG	CATATGTAAA	AAAAAAAAA	AAGGGAATTG	ACATGTCACA	TATCAAATGA	1020
ATGGAAACTT	TGTTGAAACT	TTAAAAAGCA	AATTTACTCC	AAAGACTTGT	ATTGGAAATT	1080
ACATACCTTT	TTTTTTTTT	TTTAAAGGAC	TACAGATTAT	TTTTAATGAC	TAAATTGGAG	1140
TGATACTTCT	TACACTAAAA	ATTATTTCTT	AGGCATTCTG	AATCTGGGAT	GAGAAACAGG	1200
ATTGTTTCAC	AATAGTAAGC	ACATAATTTT	TAAGGCCAAG	GCACATTTGA	CTCCTGAGAT	1260
GAATTTTTTG	TGGTCATAAT	CAAATACTTA	GTTGTTTTTG	ATGCCCCAAA	ATAAAGTGAG	1320
AATGGTAATT	TGCCAGGAAT	TCTTCATAAC	AGTATCTTAC	AAAAAACGTG	TTGCTCTCTT	1380
CACAGTATTA	TGTGTAAAGT	CATTGTTTAA	AGCACGAATG	TTCCCTCTGG	GGTACTTGTT	1440
AAAGCTAAAT	TTATTTTGCT	TCCCTCCACT	TAGAAGTGCT	GCACACTTTA	CAGCAGCTTC	1500
CTTTCTTTCC	ATGGCACTGC	CTAGTTAACA	GAAGTCTTAT	AAAAATTTAA	AAAGACACAT	1560
TTCTTACAAA	AAAGAGTTGA	ATGAGGTAAA	ATGGCATTAG	ATGGCTCTAT	ATTTTTTAAA	1620
GCTATGTAAT	TGTTCAGCGT	CACTTTTCTA	AGTACTTATA	CATATCTAAA	CATGTCTTCA	1680
TGGTTTATAT	TTTCACTTAT	ATATGCTGGG	CTGGATTAAG	CTTTGTTGTG	ATTGTGACCA	1740
ACATTCAGGC	CACGTGAGCA	CTGTCTTATC	ACATCGCCAA	TTAGTTGTAA	TAAACGTTCA	1800
ACGTACAAAC	ACTGGAGTGT	GTTTTTATCT	CTTTCCAAAA	GTTTGTCAAA	CTATGCAGAG	1860
CTGCTGAAGG	AAGAATTTCT	CATTTTTTT	TCAGTAAAAT	GTTGAAAATT	CCCCTCCATT	1920
TGAATATGGT	GGTTGTTATA	AGCACACA	AGATACATGG	TGGAAGATCT	AG	1972

# (2) INFORMATION FOR SEQ ID NO: 138:

# (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1741 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
  (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: THP1AZS08 (B) CLONE: 2753496
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 138:

CGGGTTCCGG	GCTCCGGGCT	CTGGGTGGCG	GCGGCTGTGA	GCNGCGGCTG	ANCCNCCGCG	60
CTGCGCANCG	ACGCGGGAAT	GAAGCGGGCG	CTGGGCAGGC	GAAAGGGCGT	GTGGTTGCGC	120
CTGAGGAAGA	TACTTTTCTG	TGTTTTGGGG	TTGTACATTG	CCATTCCATT	TCTCATCAAA	180
CTATGTCCTG	GAATACAGGC	CAAACTGATT	TTCTTGAATT	TCGTAAGAGT	TCCCTATTTC	240
ATTGATTTGA	AAAAACCACA	GGATCAAGGT	TTGAATCACA	CGTGTAACTA	CTACCTGCAG	300
CCAGAGGAAG	ACGTGACCAT	TGGAGTCTGG	CACACCGTCC	CTGCAGTCTG	GTGGAAGAAC	360
GCCCAAGGCA	AAGACCAGAT	GTGGTATGAG	GATGCCTTGG	CTTCCAGCCA	CCCTATCATT	420
CTGTACCTGC	ATGGGAACGC	AGGTACCAGA	GGAGGCGACC	ACCGCGTGGA	GCTTTACAAG	480
GTGCTGAGTT	CCCTTGGTTA	CCATGTGGTC	ACCTTTGACT	ACAGAGGTTG	GGGTGACTCA	540
GTGGGAACGC	CATCTGAGCG	GGGCATGACC	TATGACGCAC	TCCACGTTTT	TGACTGGATC	600
AAAGCAAGAA	GTGGTGACAA	CCCCGTGTAC	ATCTGGGGCC	ACTCTCTGGG	CACTGGCGTG	660
GCGACAAATC	TGGTGCGGCG	CCTCTGTGAG	CGAGAGACGC	CTCCAGATGC	CCTTATATTG	720
GAATCTCCAT	TCACTAATAT	CCGTGAAGAA	GCTAAGAGCC	ATCCATTTTC	AGTGATATAT	780
CGATACTTCC	CTGGGTTTGA	CTGGTTCTTC	CTTGATCCTA	TTACAAGTAG	TGGAATTAAA	840
TTTGCAAATG	ATGAAAACGT	GAAGCACATC	TCCTGTCCCC	TGCTCATCCT	GCACGCTGAG	900
GACGACCCGG	TGGTGCCCTT	CCAGCTTGGC	AGAAAGCTCT	ATAGCATCGC	CGCACCAGCT	960
CGAAGCTTCC	GAGATTTCAA	AGTTCAGTTT	GTGCCCTTTC	ATTCAGACCT	TGGCTACAGG	1020
CACAAATACA	TTTACAAGAG	CCCTGAGCTG	CCACGGATAC	TGAGGGAATT	CCTGGGGAAG	1080
TCGGAGCCTG	AGCACCAGCA	CTGAGCCTGG	CCGTGGGAAG	GAAGCATGAA	GACCTCTGCC	1140
CTCCTCCCGT	TTTCCTCCAG	TCAGCAGCCC	GGTATCCTGA	AGCCCCGGGG	GGCCGGCACC	1200
TGCAATGCTC	AGGAGCCCAG	CTCGCACCTG	GAGAGCACCT	CAGATCCCAG	GTGGGGAGGC	1260
CCCTGCAGGC	CTGCAGTGCC	CGGAGGCCTG	AGCATGGCTG	TGTGGAAAGC	GTGGGTGGCA	1320
GGCATGTGGC	TCTCCTTGCC	GCCCCTCAAC	CTGAGATCTT	GTTGGGAGAC	TTAATGGCAG	1380
CAGGCAGCCA	TCACTGCCTG	GTTGATGCTG	CACTGAGCTG	GACAGGGGGA	GTCCGGGCAG	1440
GGGACTCTTG	GGGCTCGGGA	CCATGCTGAG	CTTTTTGGCA	CCACCCACAG	AGAACGTGGG	1500

GTCCAGGTTC TTTCTGCACC TTCCCAGCAC ATGCAGAATG ACTCCAGTGG TTCCATCGTC 1560 CCCTCCTGCC CTGTGTACCT GCTTGCCTTT CTCAGCTGCC CCACCTCCCC TGGGCTGGCC 1620 CACTCACCCA CAGTGGAAGT GCCCGGGATC TGCACTTCCT CCCCTTTCAC CTACCTGTAC 1680 ACCTAACCTG GCCTTAGACT GAGCTTTATT TAAGAATAAA ATCGTGGTGG TGAAAAAAAA 1740 1741 Α

# (2) INFORMATION FOR SEQ ID NO: 139:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2808 base pairs

  - (B) TYPE: nucleic acid(C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: OVARTUT03
- (B) CLONE: 2781553

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 139:

GGCAAGATGG	CGGAAGGGGA	GGACGTGGGA	TGGTGGCGGA	GCTGGCTGCA	GCAGAGCTAC	60
CAAGCAGTCA	AAGAGAAGTC	CTCTGAAGCC	TTGGAGTTTA	TGAAGCGGGA	CCTGACGGAG	120
TTTACCCAGG	TGGTGCAGCA	TGACACGGCC	TGTACCATCG	CAGCCACGGC	CAGCGTGGTC	180
AAGGAGAAGC	TGGCTACGGA	AGGCTCCTCA	GGAGCAACAG	AGAAGATGAA	GAAAGGGTTA	240
TCTGACTTCC	TAGGGGTGAT	CTCAGACACC	TTTGCCCCTT	CGCCAGACAA	AACCATCGAC	300
TGCGATGTCA	TCACCCTGAT	GGGCACACCG	TCTGGCACAG	CTGAGCCCTA	TGATGGCACC	360
AAGGCTCGCC	TCTATAGCCT	GCAGTCGGAC	CCAGCAACCT	ACTGTAATGA	ACCAGATGGG	420
CCCCGGAAT	TGTTTGACGC	CTGGCTTTCC	CAGTTCTGCT	TGGAGGAGAA	GAAGGGGGAG	480
ATCTCAGAGC	TCCTTGTAGG	CAGCCCCTCC	ATCCGGGCCC	TCTACACCAA	GATGGTTCCA	540
GCAGCTGTTT	CCCATTCAGA	ATTCTGGCAT	CGGTATTTCT	ATAAAGTCCA	TCAGTTAGAG	600
CAGGAGCAGG	CCCGGAGGGA	CGCCCTGAAG	CAGCGGGCGG	AACAGAGCAT	CTCTGAAGAG	660
CCCGGCTGGG	AGGAGGAGGA	AGAGGAGCTC	ATGGGCATTT	CACCCATATC	TCCAAAAGAG	720
GCAAAGGTTC	CTGTGGCCAA	AATTTCTACA	TTCCCTGAAG	GAGAACCTGG	CCCCCAGAGC	780
CCCTGTGAAG	AGAATCTGGT	GACTTCAGTT	GAGCCCCCAG	CAGAGGTGAC	TCCATCAGAG	840
AGCAGTGAGA	GCATCTCCCT	CGTGACACAG	ATCGCCAACC	CGGCCACTGC	ACCTGAGGCA	900
CGAGTGCTAC	CCAAGGACCT	GTCCCAAAAG	CTGCTAGAGG	CATCCTTGGA	GGAACAGGGC	960
CTGGCTGTGG	ATGTGGGTGA	GACTGGACCC	TCACCCCCTA	TTCACTCCAA	GCCCCTAACG	1020

CCTGCTGGCC	ACACCGGCGG	CCCAGAGCCC	AGGCCTCCAG	CCAGAGTAGA	GACTCTGAGG	1080
GAGGAGGCGC	CCACAGACTT	ACGGGTGTTT	GAGCTGAACT	CGGATAGTGG	GAAGTCTACA	1140
CCCTCCAACA	ATGGAAAGAA	AGGCTCAAGC	ACGGACATCA	GTGAGGACTG	GGAGAAAGAC	1200
TTTGACTTGG	ACATGACTGA	AGAGGAGGTG	CAGATGGCAC	TTTCCAAAGT	GGATGCCTCC	1260
GGGGAGCTGG	AAGATGTAGA	GTGGGAGGAC	TGGGAGTGAG	GGAGCCAGAG	GGAGCAGCTC	1320
CCCCACCCAT	GGCATCTCTC	GCCTCCCTCG	CTCGTCTCAG	CCCAGCCCTG	GAAGACTGAG	1380
AATGTTCCCC	CAAATCTCCT	CTGCCAACCA	GAGCTCTGGG	CACAGATTCT	GGTGGCTCCC	1440
TGCTGGCCCT	CTTGGGCCTC	TGCTCACACC	TGGGAAGGGG	CTCTCTAAAT	CCCGGCCAGA	1500
AACTCTGACT	TGTGCCAACA	ATAGGATGAC	CCAAGGGAGA	GGAAACCTAT	CCTCCTCACC	1560
AGAAGAGCCT	GTGTTTTCT	GCTGAACACC	CACTGTTCCT	GAGGACTCCT	GCTGGGAAGT	1620
CCCAAGGGAT	AGTTCTAGCC	CTTCTGCCTG	TGTAGACAGA	AGCTAAACCA	CCAGTCTCTC	1680
TCGGAGGAAG	CTGAGACAAC	ATACTCTGTC	CATACATAAG	CAGGCAGGGA	GGGCCATGCC	1740
ACCTACCCTT	GGCTAAACAG	GGACAGTGAA	CACATTTTGG	TTCCTATCCC	AGTGGGTAAG	1800
AGGCACTTAT	CTCTGGGAAA	TTTGCCTCTC	TTGGGACTCT	CCCCCTCCCA	GGCATTTTCC	1860
ATTCCTGGAA	AGGCTCCTTT	GGGGTTCAGA	ATCCAGAGAC	CAAACCCTGA	CCCACCTCCT	1920
TCCTTTCCTC	CAGCCCACGC	TGGTCTGTCC	CCATGCCTTC	CCAGGGCTTC	TTCATGTCAG	1980
ATGCACCCAA	GTCCTTAGCC	CAGCTGTGCC	ACCTGCAGGA	GTTCGCTCTT	GCGTTTCTTC	2040
CCCTCCCCAA	GAAGGGAGGG	GGCTACTTCA	GGCCCTTCTG	TGTGTTGCCT	GGCAGGATAC	2100
CTTGTCCAAC	CAGCTACCCA	CCTCAACTCC	CCTGTAGTTT	AGGACACAAA	ACAGCTACCA	2160
GCGGTACAGA	GCGGTGATCA	AAGCCGAGTA	CTTACAACTC	TGGTAAGCCT	AGCTTCTCCG	2220
CCTCAGCCCT	TCTGCTTCTG	GAAGGGCTAT	CCTGGGGGTG	AACTTGAAAC	TCTCATCAGG	2280
CTTCTGCAAA	AGCTCTTCTT	CCTGAAGACA	GACCCAGCCT	TTGTGCTCTC	ACCCTCCACT	2340
CTGGTAAAGC	TGCACCTCTG	GGGGAATGAG	GGGCTGCAGG	AATCTCTGGA	GAGCCTGGTG	2400
CTTCACGATG	CTGCTCTGGT	GATTCTTGTA	CCTAATCTGG	TGTGCTCACC	AATGAGTGAA	2460
AGGGATCGTG	GGTCAGGGAC	ACCGAGAGAG	TGAGGTCACT	TCCACTTCAA	ACCTTCAGTG	2520
AGGGGGTGGG	ATGGAGAGAA	TGCTGAATCT	TTTTTTTGAC	GGGATGGGGT	TTTTCTCTTT	2580
GTAATTATTT	CTTTAGTTTA	ATTAACCTTT	TGGTTGTTTG	TGCAATATTA	TATATTTTAA	2640
ATTATAATGC	ATCTCCCCAG	AGTATTTTGT	AGCTGGGAAA	AGAAAAAAGG	AAAAAAAGAA	2700
AAAAAGATTC	TAACAGCTGT	TAGTTTTATA	ATTAAAAAAG	AAAGAAAAA	GAACTTTGTC	2760
CTGAACCTTT	TACAGACTTG	CCGTTAACAG	CATTAAAGTG	ATTCACCC		2808

- (2) INFORMATION FOR SEQ ID NO: 140:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 717 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: ADRETUT06
    - (B) CLONE: 2821925
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 140:
- CATGCGCCGA CCTTCCTCGG CTGGATTTAC ANGTTNNCCC TTAACACCCG GGATTTAAGG 60
  GACCCACACT ACCTTCCCGA AGTTGAAGGC AAGCGGTGAT TGTTTGTAGA CGGCGCTTTG 120
  TCATGGGACC TGTGCAGGTG GGAATATTGC TTTTCCTTTT TTTGGCCGTG CACGAGGCTT 180
  GGGCTGGGAT GTTGAAGGAG GAGGACGATG ACACAGAACG CTTGCCCAGC AAATGCGAAG 240
  TGTGTAAGCT GCTGAGCACA GAGCTACAGG CGGAACTGAG TCGCACCGGT CGATCTCGAG 300
  AGGTGCTGGA GCTGGGGCAG GTGCTGGATA CAGGCAAGAG GAAGAGACAC GTGCCTTACA 360
  GCGTTTCAGA GACAAGGCTG GAAGAGGCCT TAGAGAATTT ATGTGAGCGG ATCCTGGACC 420
  ATAGTGTTCA CGCTGAGCGC AAGGGCTCAC TGAGATATGC CAAGGGTCAG AGTCAGACCA 480
  TGGCAACACT GAAAGGCCTA GTGCAGAGG GGGTGAAGGT GGATCTGGGG ATCCCTCTG 540
  AGCTTTTGGG ATGAGCCCAG CCGTTGAGGT CACCATCACG AGCAGCCAT 600
  GTTNGAGGAG TTTTGAGACA TTGTGGGAGA CTGGTACTTG CACCATCAGG AGCAGCCGT 660
  ACAAGATTTT CTCTGTGAAG GTCATGTGCT GCCAGCTGCT TGAACTGCAT GTCGGGT 717
- (2) INFORMATION FOR SEQ ID NO: 141:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 2552 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: UTRSTUT05
    - (B) CLONE: 2879068
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 141:

GGCAGGGGC GCGCCGGGC CAGCGCCACG TCACCGCCCA GCAGCCCTCC CGATTGGCGG 60
GCGGGGCGGC TATAAAGGGA GGGCGCAGGC GGCGCCCGGA TCTCTTCCGC CGCCATTTTA 120

AATCCAGCTC	CATACAACGC	TCCGCCGCCG	CTGCTGCCGC	GACCCGGACT	GCGCGCCAGC	180
					0000001100	240
		GTCACTATGG				
					GGTAAAATGT	
TTATTGGAGG	CTTGAGCTGG	GATACAAGCA	AAAAAGATCT	GACAGAGTAC	TTGTCTCGAT	360
TTGGGGAAGT	TGTAGACTGC	ACAATTAAAA	CAGATCCAGT	CACTGGGAGA	TCAAGAGGAT	420
TTGGATTTGT	GCTTTTCAAA	GATGCTGCTA	GTGTTGATAA	GGTTTTGGAA	CTGAAAGAAC	480
ACAAACTGGA	TGGCAAATTG	ATAGATCCCA	AAAGGGCCAA	AGCTTTAAAA	GGGAAAGAAC	540
CTCCCAAAAA	GGTTTTTGTG	GGTGGATTGA	GCCCGGATAC	TTCTGAAGAA	CAAATTAAAG	600
AATATTTTGG	AGCCTTTGGA	GAGATTGAAA	ATATTGAACT	TCCCATGGAT	ACAAAAACAA	660
ATGAAAGAAG	AGGATTTTGT	TTTATCACAT	ATACTGATGA	AGAGCCAGTA	AAAAAATTGT	720
TAGAAAGCAG	ATACCATCAA	ATTGGTTCTG	GGAAGTGTGA	AATCAAAGTT	GCACAACCCA	780
AAGAGGTATA	TAGGCAGCAA	CAGCAACAAC	AAAAAGGTGG	AAGAGGTGCT	GCAGCTGGTG	840
GACGAGGTGG	TACGAGGGGT	CGTGGCCGAG	GTCAGGGCCA	AAACTGGAAC	CAAGGATTTA	900
ATAACTATTA	TGATCAAGGA	TATGGAAATT	ACAATAGTGC	CTATGGTGGT	GATCAAAACT	960
ATAGTGGCTA	TGGCGGATAT	GATTATACTG	GGTATAACTA	TGGGAACTAT	GGATATGGAC	1020
AGGGATATGC	AGACTACAGT	GGCCAACAGA	GCACTTATGG	CAAGGCATCT	CGAGGGGGTG	1080
GCAATCACCA	AAACAATTAC	CAGCCATACT	AAAGGAGAAC	ATTGGAGAAA	ACAGGTGTGT	1140
ATAAGAGTAC	AGGAAAACAG	TAGAAATGTC	TAATTTAATT	TAAAGATCAA	TAGACAAATG	1200
AAACGTAAAA	ACAAAATACT	ATGTAGCCTG	TTTTTACTAA	ATTGTTGATT	TTTTAATTGC	1260
TTTATGAGCC	TGTTTTGCCT	AAAGTGTCTA	TAGATCTTTA	ACTTTAAAGT	CTTATCTCAC	1320
TTTCTTTAGT	ATTGCAGAAA	AACTTAAGAG	TTTTTCTGTT	TGCTTTTGTG	TACCAGGTGG	1380
TCTAGAGGAA	TAATTAAACA	TTTTAGAACT	ATTAACAGGT	AAAGTACTGA	AATGGGTACA	1440
ACTTAAGGAA	AACAAGAATG	TTGTCTTCTA	ACTCTGACAT	TATACCTTGT	TTGTACCCGC	1500
CAGCGGGAAC	TTCATTGCAG	GCCGTGTGTC	ACCCTGACCA	CGTCTATCTC	TGGGGGTCGC	1560
ACGTTGCGGG	CAGAGCGCAA	GGCATACACC	AGAAAACGCT	GTCCTGTGGT	ATGGTCTCTT	1620
CCAACTTCAT	GTACCAGCGT	AAAGATTAAA	GTGGAAAACT	TCAGACTTTG	GCTTCATTTT	1680
TAATCTTTTT	GGAGATTAAG	TGTCTAAACT	TAACTTAAAT	GGTTTTTTAC	AGGAGTTAAA	1740
GTACATAAAT	GCCTTTTTAC	AGCTTAATCA	TTTTGGTCTT	CTGTTTAGTG	TTGTATTTCA	1800
ATTGTGGAGC	CTCATTTTAA	GTGTTCATTC	TTTTAAGATT	TAATGCTTGC	TTTTTCTTTT	1860
TATAGCTAAT	AGTGAAATCT	ACAAACCAAA	ACAAGAACTT	TTAAATCTGG	GATATAAATT	1920
					AATTGGTGTT	

TGTGATAGCA TTTTACTTGG GTTACTAGAG ATGCTTCTAG TAGACCTTAA TCTAGCATAG 2040
TTGAACCTCT GAATATGGGA AGGTTGTATT CCCAGATTCT TTCCTGAATA GATTTGAATT 2100
TAATGTCATT TGGGAACTCC AGGGTGAGTT TATTGACTAC CCAAACTGTA TTTTACCAAT 2160
AAATATGCAT ATGATCTTTA ATTATTGAAG AAAATAAAGT GAGGACTTAA AACAATTCAT 2220
GAAAGTGGAC CTTTAAAAGC TTGTCAGAGT TGCACAAATC TAACTGGTAT TTTGTTTTTG 2280
TTTTTAGGAG GAGATGTTAA AGTAACCCAT CTTGCAGGAC GACATTGAAG ATTGGTCTTC 2340
CAACTGTATA TAGCTGCCAA TTAGTTTCT TTGTTTTAC TTTGTCCTTT GCTATCTGT 2460
TTATGACTCA ATGTGGATTT GTTATACAC ATTTTATTTG TATCATTTCA TGTTAAACCC 2520
CAAATAAATG CTTCCTTATG TGAAAAAAAA AA

# (2) INFORMATION FOR SEQ ID NO: 142:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1046 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: SINJNOT02
  - (B) CLONE: 2886757
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 142:
- TACCAGTGTA AAGCCAGAGC TGAGGTTCTT GATAGTCCAC AATGGGTGAA CCACAGCAAG 60
  TGAGTGCACT TCCACCACCT CCAATGCAAT ATATCAAGGA ATATACGGAT GAAAATATTC 120
  AAGAAGGCTT AGCTCCCAAG CCTCCCCCTC CAATAAAAGA CAGTTACATG ATGTTTGGCA 180
  ATCAGTTCCA ATGTGATGAT CTTATCATCC GCCCTTTGGA AAGTCAGGGC ATCGAACGGC 240
  TTCATCCTAT GCAGTTTGAT CACAAGAAAG AACTGAGAAA ACTTAATATG TCTATCCTTA 300
  AGAAACTAGA AGATCTTAAG CTGCTTTTTG TACACGTGCA TCATCTTATA AAACGAGAAG 360
  AGACCCCACCA AGCAAGAGA ACCTTGAGAG TCATGATGGA CGTCCAGAAA CGTCAACGGC 480
  TTGAAACAGC TGAGAGATTT CAAAAGCACC TGGAACGAGT AATTGAAATG ATTCAGAATT 540
  GCTTGGCTTC TTTGCCTGAT GATTTGCCTC ATTCAGAAGC AGGAATGAGA GTAAAAACTG 600
  AACCAATGGA TGCTGATGAT AGCAACAATT GTACTGGACA GAATGAACAT CAAAGAGAAA 660
  ATTCAGGTCA TAGGAGAGAT CAGATTATAG AGAAAGATGC TGCCTTGTGT GTCCTAATTG 720

# (2) INFORMATION FOR SEQ ID NO: 143:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 1864 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: SCORNOT04
- (B) CLONE: 2964329
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 143:

GCCCTGGGCT	CGCGGCGGTG	CCGCGGGGAT	GGCGGGAGCC	GGAGCTGGAG	CCGGAGCTCG	60
CGGCGGAGCG	GCGGCGGGG	TCGAGGCTCG	AGCTCGCGAT	CCACCGCCCG	CGCACCGCGC	120
ACATCCTCGC	CACCCTCGGC	CTGCGGCTCA	GCCCTCGGCC	CGCAGGATGG	ATGGCGGGTC	180
AGGGGGCCTG	GGGTCTGGGG	ACAACGCCCC	GACCACTGAG	GCTCTTTTCG	TGGCACTGGG	240
CGCGGGCGTG	ACGGCGCTCA	GCCATCCCCT	GCTCTACGTG	AAGCTGCTCA	TCCAGGTGGG	300
TCATGAGCCG	ATGCCCCCCA	CCCTTGGGAC	CAATGTGCTG	GGGAGGAAGG	TCCTCTATCT	360
GCCGAGCTTC	TTCACCTACG	CCAAGTACAT	CGTGCAAGTG	GATGGTAAGA	TAGGGCTGTT	420
CCGAGGCCTG	AGTCCCCGGC	TGATGTCCAA	CGCCCTCTCT	ACTGTGACTC	GGGGTAGCAT	480
GAAGAAGGTT	TTCCCTCCAG	ATGAGATTGA	GCAGGTTTCC	AACAAGGATG	ATATGAAGAC	540
TTCCCTGAAG	AAAGTTGTGA	AGGAGACCTC	CTACGAGATG	ATGATGCAGT	GTGTGTCCCG	600
CATGTTGGCC	CACCCCTGC	ATGTCATCTC	AATGCGCTGC	ATGGTCCAGT	TTGTGGGACG	660
GGAGGCCAAG	TACAGTGGTG	TGCTGAGCTC	CATTGGGAAG	ATTTTCAAAG	AGGAAGGGCT	720
GCTGGGATTC	TTCGTTGGAT	TAATCCCTCA	CCTCCTGGGC	GATGTGGTTT	TCTTGTGGGG	780
CTGTAACCTG	CTGGCCCACT	TCATCAATGC	CTACCTGGTG	GATGACAGCT	TCAGCCAGGC	840
CCTGGCCATC	CGGAGCTATA	CCAAGTTCGT	GATGGGGATT	GCAGTGAGCA	TGCTGACCTA	900
CCCCTTCCTG	CTAGTTGGCG	ACCTCATGGC	TGTGAACAAC	TGCGGGCTGC	AAGCTGGGCT	960

# CCCCCCTTAC TCCCCAGTGT TCAAATCCTG GATTCACTGC TGGAAGTACC TGAGTGTGCA 1020 GGGCCAGCTC TTCCGAGGCT CCAGCCTGCT TTTCCGCCGG GTGTCATCAG GATCGTGCTT 1080 TGCCCTGGAG TAACCTGAAT CATCTAAAAA ACACGGTCTC AACCTGGCCA CCGTGGGTGA 1140

GGCCTGACCA CCTTGGGACA CCTGCGAGAC GACTCCAACC CAACAACAAC CAGATGTGCT 1200

GGCCTGACCA CCTTGGGACA CCTGCGAGAC GACTCCAACC CAACAACAAC CAGATGTGCT 1200

CCAGCCCAGC CGGGCTTCAG TTCCATATTT GCCATGTGTC TGTCCAGATG TGGGGTTGAG 1260

CGGGGGTGGG GCTGCACCCA GTGGATTGGG TCACCCGGCA GACCTAGGGA AGGTGAGGCG 1320

AGGTGGGGAG TTGGCAGAAT CCCCATACCT CGCAGATTTG CTGAGTCTGT CTTGTGCAGA 1380

GGGCCAGAGA ATGGCTTATG GGGGCCCAGG TTGGATGGGG AAAGGCTAAT GGGGTCAGAC 1440

CCCACCCGT CTACCCCTCC AGTCAGCCCA GCGCCCATCC TGCAGCTCAG CTGGGAGCAT 1500

CATTCTCCTG CTTTGTACAT AGGGTGTGGT CCCCTGGCAC GTGGCCACCA TCATGTCTAG 1560

GCCTATGCTA GGAGGCAAAT GGCCAGGCTC TGCCTGTGTT TTTCTCAACA CTACTTTTCT 1620

GATATGAGGG CAGCACCTGC CTCTGAATGG GAAATCATGC AACTACTCAG AATGTGTCCT 1680

CCTCATCTAA TGCTCATCTG TTTAATGGTG ATGCCTCGCG TACAGGATCT GGTTACCTGT 1740

GCAGTTGTGA ATACCCAGAG GTTGGGCAGA TCAGTGTCTC TAGTCCTACC CAGTTTTAAA 1800

GTTCATGGTA AGATTTGACC TCATCTCCCG CAAATAAATG TATTGGTGAT TTGGAAAAAA 1860

AAAA 1864

# (2) INFORMATION FOR SEQ ID NO: 144:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2295 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: SCORNOT04
- (B) CLONE: 2965248

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 144:

GTCTGCAGCT CCGGCCGCCA CTTGCGCCTC TCCAGCCTCC GCAGGCCCAA CCGCCGCCAG 60

CACCATGGCC AGCACCATTT CCGCCTACAA GGAGAAGATG AAGGAGCTGT CGGTGCTGTC 120

GCTCATCTGC TCCTGCTTCT ACACACAGCC GCACCCCAAT ACCGTCTACC AGTACGGGGA 180

CATGGAGGTG AAGCAGCTGG ACAAGCGGGC CTCAGGCCAG AGCTTCGAGG TCATCCTCAA 240

GTCCCCTTCT GACCTGTCCC CAGAGAGCCC TATGCTCTCC TCCCCACCCA AGAAGAAGGA 300

CACCTCCCTG GAGGAGCTGC AAAAGCGGCT GGAGGCAGCC GAGGAGCGGA GGAAGACGCA 360

GGAGGCGCAG	GTGCTGAAGC	AGCTGGCGGA	CGGCGCGAGC	ACGAGCGCGA	GGTGCTGCAC	420
AAGGCGCTGG	AGGAGAATAA	CAACTTCAGC	CGCCAGGCGG	AGGAGAAGCT	CAACTACAAG	480
ATGGAGCTCA	GCAAGGAGAT	CCGCGAGGCA	CACCTGGCCG	CACTGCGCGA	GCGGCTGCGC	540
GAGAAGGAGC	TGCACGCGGC	CGAGGTGCGC	AGGAACAAGG	AGCAGCGAGA	AGAGATGTCG	600
GGCTAAGGGC	CCGGGACGGG	CGGCGCCCAT	CCTGCGACGG	AACACGTTCG	GGTTTTGGTT	660
TTGTTTCGTT	CACCTCTGTC	TAGATGCAAC	TTTTGTTCCT	CCTCCCCCAC	CCCAGCCCCC	720
AGCTTCATGC	TTCTCTTCCG	CACTCAGCCG	CCCTGCCCTG	TCCTCGTGGT	GAGTCGCTGA	780
CCACGGCTTC	CCCTGCAGGA	GCCGCCGGGC	GTGAGACGCG	GTCCCTCGGT	GCAGACACCA	840
GGCCGGGCGC	GGCTGGGTCC	CCCGGGGGCC	CTGTGAGAGA	GGTGGTGGTG	ACCGTGGTAA	900
ACCCAGGGCG	GTGGCGTGGG	ATCGCGGGTC	CTTACGCTGG	GCTGTCTGGT	CAGCACGTGC	960
AGGTCAGGGC	AGGTCCTCTG	AGCCGGCGCC	CCTGGCCAGC	AGGCGAGGCT	ACAGTACCTG	1020
CTGTCTTTCC	AGGGGGAAGG	GGCTCCCCAT	GAGGGAGGG	CGACGGGGA	GGGGGGTGAT	1080
GGTGCCTGGG	AGCCTGCGTG	TGCAGCCGGT	GCTTGTTGAA	CTGGCAGGCG	GGTGGGTGGG	1140
GGCTGCAGCT	TTCCTTAATG	TGGTTGCACA	GGGGTCCTCT	GAGACCACCT	GGCGTGAGGT	1200
GGACACCCTG	GGCCTTCCTG	GAAGCCTGCA	GTTGGGGGCC	TGCCCTGAGT	CTGCTGGGGA	1260
GTGGGCATTC	TCTGCCAGGG	ACCCATGAGC	AGGCTGCATG	GTCTAGAGGT	TGTGGGCAGC	1320
ATGGACAGTC	CCCCACTCAG	AAGTGCAAGA	GTTCCAAAGA	GCCTCTGGCC	CAGGCCCCTC	1380
CGTGGGACAG	CCCCGCCGCC	CCTCCCCACC	AGGGCTTTGC	AGATGTCCTT	GAAAGACCCA	1440
CCCTAGAGCC	CTTTGGAGTG	CTGGCCCCTC	CTGTGCCCTC	TGCCCTGGTG	GAAGCGGCAG	1500
CCACAAGTCC	TCCTCAGGGA	GCCCCAAGGG	GGATTTTGTG	GGACCGCTGC	CCACAGATCC	1560
AGGTGTTGGA	AGGGCAGCGG	GTAAGGTTCC	CAAGCCAGCC	CCAACACCCT	TCCCACTTGG	1620
CACCCAGAGG	GGGCTGTGGG	TGGAGGCCTG	ACTCCAGGCC	TCTCCTGCCC	ACACCCTCTG	1680
GGCTGAGTTC	CTTCTTTCCC	TTGGACGCCC	AGTGCTGGCC	TTGGAGGACG	GTCAGCTGGA	. 1740
GGATGGCGGT	' GGGGGAGGCT	GTCTTTGTAC	CACTGCAGCA	TCCCCCACTI	CTCCACGGAA	1800
GCCCCATCCC	: AAAGCTGCTG	CCTGGCCCCT	' TGCTGTAAAG	TGTGAAGGG	GCGGCTGAGT	1860
TCTCTTAGGA	CCCAGAGCCA	GGGCCCTCAA	CTTCCATCCT	GCGGGAGGCC	TTGGCCGGGC	1920
ACTGCCAGTG	TCTTCCAGAG	CCACACCCAG	GGACCACGGG	AGGATCCTG <i>P</i>	A CCCCTGCAGG	1980
GCTCAGGGGT	CAGCAGGGAC	CCACTGCCC	ATCTCCCTCT	CCCCACCAA	ACAGCCCCAG	2040
AAGGAGCAGC	CAGCTGGGAT	GGGAACCCAF	GGCTGTCCAC	: ATCTGGCTT	TGTGGGACTC	2100
AGAAAGGGAA	A GCAGAACTG <i>I</i>	A GGGCTGGGAT	ATTCCTCATE	GTGGCAGCG	C TCATAGCGAA	2160
AGCCTACTGT	T AATATGCACO	CATCTCATCO	C ACGTAGTAAA	GTGAACTTA	A AAATTCAATC	2220

AAATGAACAA TTAAATAAAC ACCTGTGTGT TTAAGACAAA ATAAAAATGG AGGAGAACAA 2280 2295 AAAAAAGGG GCGGT

- (2) INFORMATION FOR SEQ ID NO: 145:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 842 base pairs (B) TYPE: nucleic acid

    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: TLYMNOT06
    - (B) CLONE: 3000534
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 145:

CGGGGACGGA	AGCAGCCCCT	GGGCCCGAGG	GGCTCGAGGC	CGGGCCGGGG	CGATGTGGAG	60
CGCGGGCCGC	GGCGGGGCTG	CCTGGCCGGT	GCTGTTGGGG	CTGCTGCTGG	CGCTGTTAGT	120
GCCGGGCGGT	GGTGCCGCCA	AGACCGGTGC	GGAGCTCGTG	ACCTGCGGGT	CGGTGCTGAA	180
GCTGCTCAAT	ACGCACCACC	GCGTGCGGCT	GCACTCGCAC	GACATCAAAT	ACGGATCCGG	240
CAGCGGCCAG	CAATCGGTGA	CCGGCGTAGA	GGCGTCGGAC	GACGCCAATA	GCTACTGGCG	300
GATCCGCGGC	GGCTCGGAGG	GCGGGTGCCC	GCGCGGGTCC	CCGGTGCGCT	GCGGGCAGGC	360
GGTGAGGCTC	ACGCATGTGC	TTACGGGCAA	GAACCTGCAC	ACGCACCACT	TCCCGTCGCC	420
GCTGTCCAAC	AACCAGGAGG	TGAGTGCCTT	TGGGGAAGAC	GGCGAGGGCG	ACGACCTGGA	480
CCTATGGACA	GTGCGCTGCT	CTGGACAGCA	CTGGGAGCGT	GAGGCTGCTG	TGCGCTTCCA	540
GCATGTGGGC	ACCTCTGTGT	TCCTGTCAGT	CACGGGTGAG	CAGTATGGAA	GCCCCATCCG	600
TGGGCAGCAT	GAGGTCCACG	GCATGCCCAG	TGCCAACACG	CACAATACGT	GGAAGGCCAT	660
GGAAGGCATC	TTCATCAAGC	CTAGTGTGGA	GCCCTCTGCA	GGTCACGATG	AACTCTGAGT	720
GTGTGGATGG	ATGGGTGGAT	GGAGGGTGGC	AGGTGGGGCG	TCTGCAGGGC	CACTCTTGGC	780
AGAGACTTTG	GGTTTGTAGG	GGTCCTCAAG	TGCCTTTGTG	ATTAAAGAAT	GTTGGTCTAA	840
AA						842

- (2) INFORMATION FOR SEQ ID NO:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 2345 base pairs
    - (B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(vii) IMMEDIATE SOURCE:

(A) LIBRARY: HEAANOT01

(B) CLONE: 3046870

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 146:

GTCCCGCCC GCAGCTGCGC GCAGGCGCTC GACGAGCCGC TCGCATTCTA CGTAACGGAC 60 GGCGGAGGCT ACGTGAAGAG AGGCGCGGCG TGACTGAGCT ACGGTTCTGG CTGCGTCCTA 120 GAGGCATCCG GGGCAGTAAA ACCGCTGCGA TCGCGGAGGC GGCGGCCAGG CCGAGAGGCA 180 GGCCGGCAG GGGTGTCGGA CGCAGGGCGC TGGGCCGGGT TTCGGCTTCG GCCACAGCTT 240 TTTTTCTCAA GGTGCAATGA AAGCCTTCCA CACTTTCTGT GTTGTCCTTC TGGTGTTTGG 300 GAGTGTCTCT GAAGCCAAGT TTGATGATTT TGAGGATGAG GAGGACATAG TAGAGTATGA 360 TGATAATGAC TTCGCTGAAT TTGAGGATGT CATGGAAGAC TCTGTTACTG AATCTCCTCA 420 ACGGGTCATA ATCACTGAAG ATGATGAAGA TGAGACCACT GTGGAGTTGG AAGGGCAGGA 480 TGAAAACCAA GAAGGAGATT TTGAAGATGC AGATACCCAG GAGGGAGATA CTGAGAGTGA 540 ACCATATGAT GATGAAGAAT TTGAAGGTTA TGAAGACAAA CCAGATACTT CTTCTAGCAA 600 AAATAAAGAC CCAATAACGA TTGTTGATGT TCCTGCACAC CTCCAGAACA GCTGGGAGAG 660 TTATTATCTA GAAATTTTGA TGGTGACTGG TCTGCTTGCT TATATCATGA ATTACATCAT 720 TGGGAAGAAT AAAAACAGTC GCCTTGCACA GGCCTGGTTT AACACTCATA GGGAGCTTTT 780 GGAGAGCAAC TTTACTTTAG TGGGGGATGA TGGAACTAAC AAAGAAGCCA CAAGCACAGG 840 AAAGTTGAAC CAGGAGAATG AGCACATCTA TAACCTGTGG TGTTCTGGTC GAGTGTGCTG 900 TGAGGGCATG CTTATCCAGC TGAGGTTCCT CAAGAGACAA GACTTACTGA ATGTCCTGGC 960 CCGGATGATG AGGCCAGTGA GTGATCAAGT GCAAATAAAA GTAACCATGA ATGATGAAGA 1020 CATGGATACC TACGTATTTG CTGTTGGCAC ACGGAAAGCC TTGGTGCGAC TACAGAAAGA 1080 GATGCAGGAT TTGAGTGAGT TTTGTAGTGA TAAACCTAAG TCTGGAGCAA AGTATGGACT 1140 GCCGGACTCT TTGGCCATCC TGTCAGAGAT GGGAGAAGTC ACAGACGGAA TGATGGATAC 1200 AAAGATGGTT CACTTTCTTA CACACTATGC TGACAAGATT GAATCTGTTC ATTTTTCAGA 1260 CCAGTTCTCT GGTCCAAAAA TTATGCAAGA GGAAGGTCAG CCTTTAAAGC TACCTGACAC 1320 TAAGAGGACA CTGTTGTTTA CATTTAATGT GCCTGGCTCA GGTAACACTT ACCCAAAGGA 1380 TATGGAGGCA CTGCTACCCC TGATGAACAT GGTGATTTAT TCTATTGATA AAGCCAAAAA 1440 GTTCCGACTC AACAGAGAAG GCAAACAAAA AGCAGATAAG AACCGTGCCC GAGTAGAAGA 1500 GAACTTCTTG AAACTGACAC ATGTGCAAAG ACAGGAAGCA GCACAGTCTC GGCGGGAGGA 1560

GAAAAAAGA GCAGAGAAGG AGCGAATCAT GAATGAGGAA GATCCTGAGA AACAGCGCAG 1620
GCTGGAGGAG GCTGCATTGA GGCGTGAGCA AAAGAAGTTG GAAAAGAAGC AAATGAAAAT 1680
GAAACAAATC AAAGTGAAAG CCATGTAAAG CCATCCCAGA GATTTGAGTT CTGATGCCAC 1740
CTGTAAGCTC TGAATTCACA GGAAACATGA AAAACGCCAG TCCATTTCTC AACCTTAAAT 1800
TTCAGACAGT CTTGGGCAAC TGAGAAATCC TTATTTCATC ATCTACTCTG TTTGGGGTTT 1860
GGGGTTTTAC AGAGATTGAA GATACCTGGA AAGGGCTCTG TTTCAAGAAT TTTTTTTCC 1920
AGAATAATCAA ATTATTTTGA TTATTTTATA AAAGGAATGA TCTATGAAAT CTGTGTAGGT 1980
TTTAAATATT TTAAAAATTA TAATACAAAT CATCAGTGCT TTTAGTACTT CAGTGTTTAA 2040
AGAAATACCA TGAAATTTAT AGGTAGATAA CCAGATTGTT GCTTTTTGTT TAAACCAAGC 2100
AGTTGAAATG GCTATAAAGA CTGACTCTAA ACCAAGATTC TGCAAATAAT GATTGGAATT 2160
GCACAATAAA CATTGCTTGA TGTTTTCTTG TATGTCTACA TTAAACTAGA GAAAAAGTAA 2220
AAATTAGAAC ACTGTATGTA GTAATGAAAT TTCAGGGACC CAGAACATAA TGTAGTATAT 2280
GTTTTTAGGT GGGAGATGCT GATAACAAAA TTAATAGGAA GTCTGTAGGC ATTAGGATAC 2340
TGACA

# (2) INFORMATION FOR SEQ ID NO: 147:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2215 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: PONSAZT01
  - (B) CLONE: 3057669
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 147:
- CCCACGCGTC CGCCCACGCG TCCGTTTTCA GTAGGGATTT CCTGTGACCA GACAAGTTCA 60
  TCTGAGAGCC AGTTCTCACC ACTGGAATTC TCAGGAATGG ACCATGAGGA CATCAGTGAG 120
  TCAGTGGATG CAGCATACAA CCTCCAGGAC AGTTGCCTTA CAGACTGTGA TGTGGAAGAT 180
  GGGACTATGG ATGGCAATGA TGAGGGGCAC TCCTTTGAAC TTTGTCCTTC TGAAGCTTCT 240
  CCTTATGTAA GGTCAAGGGA GAGAACCTCC TCTTCAATAG TATTTGAAGA TTCTGGCTGT 300
  GATAATGCTT CCAGTAAAGA AGAGCCGAAA ACTAATCGAT TGCATATTGG CAACCATTGT 360
  GCTAATAAAC TAACTGCTTT CAAGCCCACC AGTAGCAAAA CTTCTTCTGA AGCTACATAG 420
  TCTATTTCTC CTCCAAGACC AACCACTTTA AGTTTAGATC TCACTAAAAA CACCACAGAA 480

AAACTCCAGC	CCAGTTCACC	AAAGGTGTAT	CTTTACATTC	AAATGCAGCT	GTGCAGAAAA	540
GAAAACCTCA	AAGACTGGAT	GAATGGACGA	TGTACCATAG	AGGAGAGAGA	GAGGAGCGTG	600
TGTCTGCACA	TCTTCCTGCA	GATCGCAGAG	GCAGTGGAGT	TTCTTCACAG	TAAAGGACTG	660
ATGCACAGGG	ACCTCAAGCC	ATCCAACATA	TTCTTTACAA	TGGATGATGT	GGTCAAGGTT	720
GGAGACTTTG	GGTTAGTGAC	TGCAATGGAC	CAGGATGAGG	AAGAGCAGAC	GGTTCTGACC	780
CCAATGCCAG	CTTATGCCAG	ACACACAGGA	CAAGTAGGGA	CCAAACTGTA	TATGAGCCCA	840
GAGCAGATTC	ATGGAAACAG	CTATTCTCAT	AAAGTGGACA	TCTTTTCTTT	AGGCCTGATT	900
CTATTTGAAT	TGCTGTATCC	ATTCAGCACT	CAGATGGAGA	GAGTCAGGAC	CTTAACTGAT	960
GTAAGAAATC	TCAAATTTCC	ACCATTATTT	ACTCAGAAAT	ATCCTTGTGA	GTACGTGATG	1020
GTTCAAGACA	TGCTCTCTCC	ATCCCCCATG	GAACGACCTG	AAGCTATAAA	CATCATTGAA	1080
AATGCTGTAT	TTGAGGACTT	GGACTTTCCA	GGAAAAACAG	TGCTCAGACA	GAGGTCTCGC	1140
TCCTTGAGTT	CATCGGGAAC	AAAACATTCA	AGACAGTCCA	ACAACTCCCA	TAGCCCTTTG	1200
CCAAGCAATT	AGCCTTAAGT	TGTGCTAGCA	ACCCTAATAG	GTGATGCAGA	TAATAGCCTA	1260
CTTCTTAGAA	TATGCCTGTC	CAAAATTGCA	GACTTGAAAA	GTTTGTTCTT	CGCTCAATTT	1320
TTTTGTGGAC	TACTTTTTT	ATATCAAATT	TAAGCTGGAT	TTGGGGGCAT	AACCTAATTT	1380
GAGCCAACTC	CTGAGTTTTG	CTATACTTAA	GGAAAGGGCT	ATCTTTGTTC	TTTGTTAGTC	1440
TCTTGAAACT	GGCTGCTGGC	CAAGCTTTAT	AGCCCTCACC	ATTTGCCTAA	GGAGGTAGCA	1500
GCAATCCCTA	ATATATATAT	ATAGTGAGAA	CTAAAATGGA	TATATTTTTA	TAATGCAGAA	1560
GAAGGAAAGT	CCCCCTGTGT	GGTAACTGTA	TTGTTCTAGA	AATATGCTTT	CTAGAGATAT	1620
GATGATTTTG	AAACTGATTT	CTAGAAAAAG	CTGACTCCAT	TTTTGTCCCT	GGCGGGTAAA	1680
TTAGGAATCT	GCACTATTTT	GGAGGACAAG	TAGCACAAAC	TGTATAACGG	TTTATGTCCG	1740
TAGTTTTATA	GTCCTATTTG	TAGCATTCAA	TAGCTTTATT	CCTTAGATGG	TTCTAGGGTG	1800
GGTTTACAGC	TTTTTGTACT	TTTACCTCCA	ATAAAGGGAA	AATGAAGCTT	TTTATGTAAA	1860
TTGGTTGAAA	GGTCTAGTTT	TGGGAGGAAA	AAAGCCGTAG	TAAGAAATGG	ATCATATATA	1920
TTACAACTAA	CTTCTTCAAC	TATGGACTTT	TTAAGCCTAA	TGAAATCTTA	AGTGTCTTAT	1980
ATGTAATCCT	GTAGGTTGGT	ACTTCCCCCA	AACTGATTAT	AGGTAACAGT	TTAATCATCT	2040
CACTTGCTAA	CATGTTTTTA	TTTTTCACTG	TAAATATGTT	TATGTTTTAT	TTATAAAAAT	2100
TCTGAAATCA	ATCCATTTGG	GTTGGTGGTG	TACAGAACAC	ACTTAAGTGT	GTTAACTTGT	2160
GACTTCTTTC	AAGTCTAAAT	GATTTAATAA	AACTTTTTT	AAATTAAAAA	AAAAA	2215

- (2) INFORMATION FOR SEQ ID NO: 148:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 1395 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (vii) IMMEDIATE SOURCE:
    - (A) LIBRARY: HEAONOTO3
    - (B) CLONE: 3088178
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 148:

GGTTGACATG	ATGAACAATC	GGTTTCGGAA	GGATATGATG	AAAAATGCTA	GTGAAAGTAA	60
ACTTTCGAAA	GACAACCTTA	AAAAGAGACT	TAAAGAAGAA	TTCCAACATG	CCATGGGAGG	120
AGTACCTGCC	TGGGCAGAGA	CTACTAAGCG	GAAAACATCT	TCAGATGATG	AAAGTGAAGA	180
GGATGAAGAT	GATTTGTTGC	AAAGGACTGG	GAATTTCATA	TCCACATCAA	CTTCTCTTCC	240
AAGAGGCATC	TTGAAGATGA	AGAACTGCCA	GCATGCGAAT	GCTGAACGTC	CTACTGTTGC	300
TCGGATCTCA	TCTGTGCAGT	TCCATCCCGG	TGCACAGATT	GTGATGGTTG	CTGGATTAGA	360
TAATGCTGTA	TCACTATTTC	AGGTTGATGG	GAAAACAAAT	CCTAAAATTC	AGAGCATCTA	420
TTTGGAAAGG	TTTCCAATCT	TTAAGGCTTG	TTTTAGTGCT	AATGGGGAAG	AAGTTTTAGC	480
CACGAGTACC	CACAGCAAGG	TTCTTTATGT	CTATGACATG	CTGGCTGGAA	AGTTAATTCC	540
TGTGCATCAA	GTGAGAGGTT	TGAAAGAGAA	GATAGTGAGG	AGCTTTGAAG	TCTCCCCAGA	600
TGGGTCCTTC	TTGCTCATAA	ATGGCATTGC	TGGATATTTG	CATTTGCTAG	CAATGAAGAC	660
CAAAGAACTG	ATTGGAAGCA	TGAAAATTAA	TGGAAGGGTT	GCAGCATCCA	CATTCTCTTC	720
AGATAGTAAG	AAAGTATACG	CCTCTTCGGG	GGATGGAGAA	GTTTATGTTT	GGGATGTGAA	780
CTCAAGGAAG	TGCCTTAACA	GATTTGTTGA	TGAAGGCAGT	TTATATGGAT	TAAGCATTGC	840
CACATCTAGG	AATGGACAGT	ATGTTGCTTG	TGGTTCTAAT	TGTGGAGTGG	TAAATATATA	900
CAATCAAGAT	TCTTGTCTCC	AAGAAACAAA	CCCAAAGCCA	ATAAAAGCTA	TAATGAACTT	960
GGTTACAGGT	GTTACTTCTC	TGACCTTCAA	TCCTACTACA	GAAATCTTGG	CAATTGCTTC	1020
AGAAAAAATG	AAAGAAGCAG	TCAGATTGGT	TCATCTTCCT	TCCTGTACAG	TATTTTCAAA	1080
CTTCCCAGTC	ATTAAAAATA	AGAATATTTC	TCATGTTCAT	ACCATGGATT	TTTCTCCGAG	1140
AAGTGGATAC	TTTGCCTTGG	GGAATGAAAA	GGGCAAGGCC	CTGATGTATA	GGTTGCACCA	1200
TTACTCAGAC	TTCTAAAGAG	ACTATTTGAA	GTCCAGTTGA	GTCACAAGAG	AAGCCTGTCT	1260
TGATATATCA	TCTCAGAAAC	TTTCCTGAAT	ATGTGATAAT	ATATGGAAAA	TGATTTATAG	1320
ATCCAGCTGT	GCTTAAGAGC	CAGTAATGTC	TTAATAAACA	TGTGGCAGCT	TTTGTTTGAA	1380
AAAAAAAAA	AAAGG					1395

# (2) INFORMATION FOR SEQ ID NO: 149:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2609 base pairs
    (B) TYPE: nucleic acid
    (C) STRANDEDNESS: single

  - (D) TOPOLOGY: linear

# (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: BRSTNOT19
- (B) CLONE: 3094321
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 149:

CCCGCCATGG	CACTGTCGCG	GGGGCTGCCC	CGGGAGCTGG	CTGAGGCGGT	GGCCGGGGGC	60
CGGGTGCTGG	TGGTGGGGGC	GGGCGGCATC	GGCTGCGAGC	TCCTCAAGAA	TCTCGTGCTC	120
ACCGGTTTCT	CCCACATCGA	CCTGATTGAT	CTGGATACTA	TTGATGTAAG	CAACCTCAAC	180
AGACAGTTTT	TGTTTCAAAA	GAAACATGTT	GGAAGATCAA	AGGCACAGGT	TGCCAAGGAA	240
AGTGTACTGC	AGTTTTACCC	GAAAGCTAAT	ATCGTTGCCT	ACCATGACAG	CATCATGAAC	300
CCTGACTATA	ATGTGGAATT	TTTCCGACAG	TTTATACTGG	TTATGAATGC	TTTAGATAAC	360
AGAGCTGCCC	GAAACCATGT	TAATAGAATG	TGCCTGGCAG	CTGATGTTCC	TCTTATTGAA	420
AGTGGAACAG	CTGGGTATCT	TGGACAAGTA	ACTACTATCA	AAAAGGGTGT	GACCGAGTGT	480
TATGAGTGTC	ATCCTAAGCC	GACCCAGAGA	ACCTTTCCTG	GCTGTACAAT	TCGTAACACA	540
CCTTCAGAAC	CTATACATTG	CATCGTTTGG	GCAAAGTACT	TGTTCAACCA	GTTGTTTGGG	600
GAAGAAGATG	CTGATCAAGA	AGTATCTCCT	GACAGAGCTG	ACCCTGAAGC	TGCCTGGGAA	660
CCAACGGAAG	CCGAAGCCAG	AGCTAGAGCA	TCTAATGAAG	ATGGTGACAT	TAAACGTATT	720
TCTACTAAGG	AATGGGCTAA	ATCAACTGGA	TATGATCCAG	TTAAACTTTT	TACCAAGCTT	780
TTTAAAGATG	ACATCAGGTA	TCTGTTGACA	ATGGACAAAC	TATGGCGGAA	AAGGAAACCT	840
CCAGTTCCGT	TGGACTGGGC	TGAAGTACAA	AGTCAAGGAG	AAGAAACGAA	TGCATCAGAT	900
CAACAGAATG	AACCCCAGTT	AGGCCTGAAA	GACCAGCAGG	TTCTAGATGT	AAAGAGCTAT	960
GCACGTCTTT	TTTCAAAGAG	CATCGAGACT	TTGAGAGTTC	ATTTAGCAGA	AAAGGGGGAT	1020
GGAGCTGAGC	TCATATGGGA	TAAGGATGAC	CCATCTGCAA	TGGATTTTGT	CACCTCTGCT	1080
GCAAACCTCA	GGATGCATAT	TTTCAGTATG	AATATGAAGA	GTAGATTTGA	TATCAAATCA	1140
ATGGCAGGGA	ACATTATTCC	TGCTATTGCT	ACTACTAATG	CAGTAATTGC	TGGGTTGATA	1200
GTATTGGAAG	GATTGAAGAT	TTTATCAGGA	AAAATAGACC	AGTGCAGAAC	AATTTTTTTG	1260
AATAAACAAC	CAAACCCAAG	AAAGAAGCTT	CTTGTGCCTT	GTGCACTGGA	TCCTCCCAAC	1320

CCCAATTGTT ATGTATGTGC CAGCAAGCCA GAGGTGACTG TGCGGCTGAA TGTCCATAAA 1380 GTGACTGTTC TCACCTTACA AGACAAGATA GTGAAAGAAA AATTTGCTAT GGTAGCACCA 1440 GATGTCCAAA TTGAAGATGG GAAAGGAACA ATCCTAATAT CTTCCGAAGA GGGAGAGACG 1500 GAAGCTAATA ATCACAAGAA GTTGTCAGAA TTTGGAATTA GAAATGGCAG CCGGCTTCAA 1560 GCAGATGACT TCCTCCAGGA CTATACTTTA TTGATCAACA TCCTTCATAG TGAAGACCTA 1620 GGAAAGGACG TTGAATTTGA AGTTGTTGGT GATGCCCCGG AAAAAGTGGG GCCCAAACAA 1680 GCTGAAGATG CTGCCAAAAG CATAACCAAT GGCAGTGATG ATGGAGCTCA GCCCTCCACC 1740 TCCACAGCTC AAGAGCAAGA TGACGTTCTC ATAGTTGATT CGGATGAAGA AGATTCTTCA 1800 AATAATGCCG ACGTCAGTGA AGAAGAGAGA AGCCGCAAGA GGAAATTAGA TGAGAAAGAG 1860 AATCTCAGTG CAAAGAGGTC ACGTATAGAA CAGAAGGAAG AGCTTGATGA TGTCATAGCA 1920 TTAGATTGAA CAGAAATGCC TCTAAACAGA ACCCTCTTAC TATTTAGTTT ATCTGGGCAG 1980 AACCAGATTG TTATGTCCTT TGTTCCAAAG GGAAAAAATT GACAGCAGTG ACTTGAAAAT 2040 GATTCTGCTC CCTTTGAAAG CATTCATTTT GCTAGAACTG TTAGACACAT TGCAGTATGC 2100 TGTATTGAAA GTAGGAATAT AGTTTTAAAA ACCCTTTGAA CAAAGTGTGT GCATAACCAG 2160 TCATGAGATA AAACAACACA ATGCATGTTG CCTTTTTAAT GTAAATACCC TTAGGTATCA 2220 TTAATAGTTT CAAAATATTG TGGTTTAGTA AAGTTGATAC CTGGTTATAA ATATTATGCC 2280 TTTATTTTG GCTAGAAGAA GAATTATTTT TAGCCTAGAT CTAACCATTT TCATACTCTT 2340 AACTGATTGA AACAGATTCA AAGAAGTATC GAGTGCTATG CATTGAAACT TGTTTTTAAA 2400 TGTTAGATGG CACTATGTAT ATTAATGTAA AACAATGTTA ATTTACTCAA GTTTTCAGTT 2460 TGTACCGCCT GGTATGTCTG TGTAAGAAGC CAATTTTTGT GTATTGTTAC AGTTTCAGGT 2520 TATTTATATT CGATGTTTTG TAAAACTCAA ATAACGACTA TACTTATGGA CCAAATAAAT 2580 GGCATCTGCA TTCTTGTTAA AAAAAAAA 2609

# (2) INFORMATION FOR SEQ ID NO: 150:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 3633 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LUNGTUT13
  - (B) CLONE: 3115936
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 150:

CCTGAGGGAT	CCACAGAGGG	TGCGGTCCTT	GGAGGGAGGA	CATGCAGTGC	CACGTGCCAT	60
GGACCAGCCA	GTGGACCCCA	TGGCCAGCAA	GGCTGCTCCT	GGGGCCAGTG	GGGTGGACAG	120
TCCCGCCCAC	GCAGGTGACT	GAGGTGCCAG	TGTGGGAATG	AAAATGCGGC	CTGTGCTCCT	180
GGGCCCATGC	GTCTCACGCT	GCCCTTCCTC	TCCAGGGAAG	CCTGTGTACC	TGCTACTTTT	240
TCCCGAACAA	TTCATGGTAA	AAACACAAAT	GGTATATGGA	CAAGATACTG	AATGTGGAAG	300
AAACCTACTT	GACAGTGTTG	GTGAAAATAG	GGCCAGGATT	TCACACCCGT	GAATGCTTTT	360
TACTGAAAAG	TATTTTGTGT	TTTTCTCCCA	GTTACAGAAT	GTCTGAAGGG	GACAGTGTGG	420
GAGAATCCGT	CCATGGGAAA	CCTTCGGTGG	TGTACAGATT	TTTCACAAGA	CTTGGACAGA	480
TTTATCAGTC	CTGGCTAGAC	AAGTCCACAC	CCTACACGGC	TGTGCGATGG	GTCGTGACAC	540
TGGGCCTGAG	CTTTGTCTAC	ATGATTCGAG	TTTACCTGCT	GCAGGGTTGG	TACATTGTGA	600
CCTATGCCTT	GGGGATCTAC	CATCTAAATC	TTTTCATAGC	TTTTCTTTCT	CCCAAAGTGG	660
ATCCTTCCTT	AATGGAAGAC	TCAGATGACG	GTCCTTCGCT	ACCCACCAAA	CAGAACGAGG	720
AATTCCGCCC	CTTCATTCGA	AGGCTCCCAG	AGTTTAAATT	TTGGCATGCG	GCTACCAAGG	780
GCATCCTTGT	GGCTATGGTC	TGTACTTTCT	TCGACGCTTT	CAACGTCCCG	GTGTTCTGGC	840
CGATTCTGGT	GATGTACTTC	ATCATGCTCT	TCTGTATCAC	GATGAAGAGG	CAAATCAAGC	900
ACATGATTAA	GTACCGGTAC	ATCCCGTTCA	CACATGGGAA	GAGAAGGTAC	AGAGGCAAGG	960
AGGATGCCGG	CAAGGCCTTC	GCCAGCTAGA	AGCGGGACTG	AGGCTGCCTC	ACGTGTTGCA	1020
AGAACAGTTT	TGAGCCATTG	TTAACAATGC	CTTTTTTCTT	CACATAAAGT	AGTTGATTAC	1080
GAGGGAGTCA	AATTTTCTTT	TTAAAAAGGA	GCTTCAATGA	TTTGTAACTG	AAATATCAGG	1140
TTCTAGAAGA	AACTGGCGCT	TAAACCAAAT	CGCATGGATT	TCTTTTTCAG	TGACGTTCAA	1200
GTGTTTCTCA	CGGATGGAAT	TCTAGTCAGC	TGCAGGCGGG	AAGCCAGGCG	GGTGGAGCCC	1260
ATGGGAGCAA	GGGCGAGTGG	CCGGTCCCCG	CTGTGCCAGG	TGGGCAGGCA	GGAGCAAGGC	1320
CTGCGAGGGA	GGAACGGGCC	GCTCCCCGCC	AGCCGCCTTC	CCCAGCAGCC	GCAGGTGGTG	1380
CCAGCCACTC	CACAGAGCCC	GAGGGATGAT	CTAGCCTGAT	TCCTGCGTGT	CCGAAAGAAC	1440
TTAACGTTTT	AAAGGTGATT	GTCAAGTAAC	TGTGTGGGGT	TCTAATGCCA	GTTTCCTAAT	1500
TCCATCTCAC	TGGAGATGTT	TAAAGTTGGC	CTCTATCCTA	ATGACTCAAA	ACTTGGTTCT	1560
TAACTACCAT	GATTGCTTTT	GAGGGCCCGG	AATTATAAAT	ATATATTATA	TTTTAATTGT	1620
TTGAGATTAT	TTTGACACAT	TTCTTTGATA	CGTAGAGTGT	TTTGTTTTTA	ATTTAAATCT	1680
GTCCTCATGC	AACCCTCCAT	GAGGGGCAGC	GAAGCTGGCA	GGGAGCAGAC	TGGCTTTGTA	1740
GGTTCAGCAC	TCGGCCCCC	ACTGCGGGAG	AGGCGGAACC	CACTTGCATG	TCAGCGTTTT	1800
TGATTCGAGA	AAAGAAATAC	TCTCAACGTT	TTACCAAGTG	ATTTTACCTC	CACCTTTACT	1860

AAAGTCTTTA	CCTAAAACAT	GGCAGTCGCT	GGACACAGGA	AAGCCCACCT	TTTGTTTGGC	1920
CTTTTCGAAA	GGTGACCCAT	ATTGCACAGC	AGAACATCAC	AGCTGTGGTC	CCAGATGAGA	1980
CACTGACATG	CGAGTGAAGG	CCTCTCCTCC	TGGGCCCCGG	GCTGCGCAGG	CTCCTCACTC	2040
TGGGCGGTGT	TTCCTGTCTC	AGAATTGACA	CGGTGAATGC	TTAGTGTCTG	GATTTTCTTG	2100
TGCCAGTGTT	TACATATCTG	ACATCGAGCT	CCTCTAAGAG	GCCACGTTCA	AGCTTGTGTG	2160
TCCCTGACCC	AAGATAGCCA	GTGCTGCTCC	CAGGTGGTAC	TTCTGGTACC	GTGTTGAGAC	2220
ACTTGGGATT	CTCAGACTGT	GGACAGGAGT	GTTTGTCATT	TTTCATACTG	TTTTCTTAAT	2280
AAGCGCTCAG	GCCTAAGGTG	TGACAGGAAG	TCGCACGCGC	TTGGCCAGAG	CACAGTGAAG	2340
CAAAGGACTG	GGTGCTGATG	GATGGAGCCA	CGGCGGCATC	TGCCCACCCG	GCCGCAGCCC	2400
CCAGTGCCTC	TCCTGGTGGT	CCTCCCAGTC	TAGAGGGTCA	CGGCCCCCC	GCCCTCCTCC	2460
GTCTCTGGCA	AGCTGACCTT	GACTAACCCA	GGAATACAGG	GTCATCCTCA	TTCCTAAGTA	2520
AGTCAAACAG	CAAGACATGG	TTTGCGCGGG	TCTTTGCCGG	AAGCCGGTCC	TGCTGGCCAG	2580
GTGTTTTACG	TCAGCAGGGA	AATGTGGCAC	ACGCCCTCGA	GGCATTTTAA	CACTGTGCTT	2640
CAGGAAATCT	CAAGTTCCAT	CTTGTGTTAG	TAACGTACCC	ACATTTTGCT	GGAGTTAGTT	2700
TATTAAAGAT	GCCTACGGTG	AACTCTCTGG	CGCAGGTTAA	ATGCAGTTTT	GAAAACCTGG	2760
AAACATCAAA	TGGAGGCGGG	AAATAGGCTG	GGGCCGAGCT	GAGGGGCTGA	ACACAGCAGT	2820
GACCGTGGGT	CAGCAGGTCG	CCTGCCCAGC	AGGCCCCCA	GGAGAGGGCT	CGGGCGCCCC	2880
TGGCAGCCCC	CATACCCCCA	GGACCTGGCT	CGTGAGTGCG	TCTGGGTCAG	GAAGAGACCT	2940
CTCTGTGCGT	CTCAGGCTGA	GATGCAGATT	TCTGTTTTCT	AAAACTGGAA	GCGACCTTGA	3000
CGTGTATTGA	AGGTGTGTGT	GCCAAATGCT	TCCGACGGAG	GTGCTGGCCT	TGGTTGGTTT	3060
CTCTCTGCCC	CGTGTGGTCA	TCAAGTCCTG	GGGGATGTGC	TCTGCCCAGC	CGCCCTCGGG	3120
GAGAGCAGCG	CCGCCTCCCA	TGGGGCCGTG	GGGCTGCTGT	TCTCACTGCA	CTGGCTGAAG	3180
CAACCCGCCA	GCCTCCGTGC	CCCACCCCAC	CCAGCACGCA	CTCATTCAGT	CCATTGCCTT	3240
AACACAAGCC	TGATGGGGCT	GTTTTCTCAC	AATATAAACG	AATAAAGTGT	CTTCTGGCCT	3300
ACTTCTGAAT	TACTTCTCAA	CTGTATGGTT	TGGGGAAGGG	AGGGAAACCT	AAAATCCCGT	3360
CCAAATAAGT	GAAATTCCTG	AAGAAGTGGC	TGAGTCCTAC	CAGGTTGGGG	TTAGGGAAAT	3420
GTTCTGGGTT	CAGGCGCCCC	TCCCAGGGCT	GAGAAAGCGC	AGCCAGGGAC	AGCTTTCTGT	3480
TCTCTCCCAG	GGTGGCTAGG	TTAGTATCTT	ACATGACAAA	AAACTGAGAG	TGTTCTAACT	3540
TCTGTGCAAG	CAAGGTTAAT	CCTGAGACTA	AATCTTGGCG	TTCAGACTCC	CGTAGAGGTC	3600
ATCTGTGTCC	AGGCCCACCC	GGGCGCCGGC	TCA			3633

## (2) INFORMATION FOR SEQ ID NO: 151:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2018 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LUNGTUT13
  - (B) CLONE: 3116522
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 151:

TGGCTCGCTG	GCCGCTCCTG	GAGGCGGCGG	CGGGAGCGCA	GGGGGCGCGC	GGCCCGGGGA	60
CTCGCATTCC	CCGGTTCCCC	CTCCACCCCA	CGCGGCCTGG	ACCATGGACG	CCAGATGGTG	120
GGCAGTGGTG	GTGCTGGCTG	CGTTCCCCTC	CCTAGGGGCA	GGTGGGGAGA	CTCCCGAAGC	180
CCCTCCGGAG	TCATGGACCC	AGCTATGGTT	CTTCCGATTT	GTGGTGAATG	CTGCTGGCTA	240
TGCCAGCTTT	ATGGTACCTG	GCTACCTCCT	GGTGCAGTAC	TTCAGGCGGA	AGAACTACCT	300
GGAGACCGGT	AGGGGCCTCT	GCTTTCCCCT	GGTGAAAGCT	TGTGTGTTTG	GCAATGAGCC	360
CAAGGCCTCT	GATGAGGTTC	CCCTGGCGCC	CCGAACAGAG	GCGGCAGAGA	CCACCCGAT	420
GTGGCAGGCC	CTGAAGCTGC	TCTTCTGTGC	CACAGGGCTC	CAGGTGTCTT	ATCTGACTTG	480
GGGTGTGCTG	CAGGAAAGAG	TGATGACCCG	CAGCTATGGG	GCCACAGCCA	CATCACCGGG	540
TGAGCGCTTT	ACGGACTCGC	AGTTCCTGGT	GCTAATGAAC	CGAGTGCTGG	CACTGATTGT	600
GGCTGGCCTC	TCCTGTGTTC	TCTGCAAGCA	GCCCCGGCAT	GGGGCACCCA	TGTACCGGTA	660
CTCCTTTGCC	AGCCTGTCCA	ATGTGCTTAG	CAGCTGGTGC	CAATACGAAG	CTCTTAAGTT	720
CGTCAGCTTC	CCCACCCAGG	TGCTGGCCAA	GGCCTCTAAG	GTGATCCCTG	TCATGCTGAT	780
GGGAAAGCTT	GTGTCTCGGC	GCAGCTACGA	ACACTGGGAG	TACCTGACAG	CCACCCTCAT	840
CTCCATTGGG	GTCAGCATGT	TTCTGCTATC	CAGCGGACCA	GAGCCCCGCA	GCTCCCCAGC	900
CACCACACTC	TCAGGCCTCA	TCTTACTGGC	AGGTTATATT	GCTTTTGACA	GCTTCACCTC	960
AAACTGGCAG	GATGCCCTGT	TTGCCTATAA	GATGTCATCG	GTGCAGATGA	TGTTTGGGGT	1020
CAATTTCTTC	TCCTGCCTCT	TCACAGTGGG	CTCACTGCTA	GAACAGGGGG	CCCTACTGGA	1080
GGGAACCCGC	TTCATGGGGC	GACACAGTGA	GTTTGCTGCC	CATGCCCTGC	TACTCTCCAT	1140
CTGCTCCGCA	TGTGGCCAGC	TCTTCATCTT	TTACACCATT	GGGCAGTTTG	GGGCTGCCGT	1200
CTTCACCATC	ATCATGACCC	TCCGCCAGGC	CTTTGCCATC	CTTCTTTCCT	GCCTTCTCTA	1260
TGGCCACACT	GTCACTGTGG	TGGGAGGGCT	GGGGGTGGCT	GTGGTCTTTG	CTGCCCTCCT	1320
GCTCAGAGTC	TACGCGCGGG	GCCGTCTAAA	GCAACGGGGA	AAGAAGGCTG	TGCCTGTTGA	1380

GTCTCCTGTG	CAGAAGGTTT	GAGGGTGGAA	AGGGCCTGAG	GGGTGAAGTG	AAATAGGACC	1440
CTCCCACCAT	CCCCTTCTGC	TGTAACCTCT	GAGGGAGCTG	GCTGAAAGGG	CAAAATGCAG	1500
GTGTTTTCTC	AGTATCACAG	ACCAGCTCTG	CAGCAGGGGA	TTGGGGAGCC	CAGGAGGCAG	1560
CCTTCCCTTT	TGCCTTAAGT	CACCCATCTT	CCAGTAAGCA	GTTTATTCTG	AGCCCCGGGG	1620
GTAGACAGTC	CTCAGTGAGG	GGTTTTGGGG	AGTTTGGGGT	CAAGAGAGCA	TAGGTAGGTT	1680
CCACAGTTAC	TCTTCCCACA	AGTTCCCTTA	AGTCTTGCCC	TAGCTGTGCT	CTGCCACCTT	1740
CCAGACTCAC	TCCCCTCTGC	AAATACCTGC	ATTTCTTACC	CTGGTGAGAA	AAGCACAAGC	1800
GGTGTAGGCT	CCAATGCTGC	TTTCCCAGGA	GGGTGAAGAT	GGTGCTGTGC	TGAGGAAAGG	1860
GGATGCAGAG	CCCTGCCCAG	CACCACCACC	TCCTATGCTC	CTGGATCCCT	AGGCTCTGTT	1920
CCATGAGCCT	GTTGCAGGTT	TTGGTACTTT	AGAAATGTAA	CTTTTTGCTC	TTATAATTTT	1980
AATTATTTA	ATTAAATTAC	TGCAGTGGAA	AAAAAAA			2018

#### (2) INFORMATION FOR SEQ ID NO: 152:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 942 base pairs

  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LUNGTUT13
  - (B) CLONE: 3117184
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 152:
- CCTCCATCAG CTCGCCGCG AGCGGCTGTA TTTGCGGCCT GTGCGAGTAG GCGCTTGGGC 60 ACTCAGTCTC CCTGGCGGGC GACGGGCAGA AATCTCGAAC CAGTGGAGCG CACTCGTAAC 120 CTGGATCCCA GAAGGTCGCG AAGGCAGTAC CGTTTCCTCA GCGGCGGACT GCTGCAGTAA 180 GAATGTCTTT TCCACCTCAT TTGAATCGCC CTCCCATGGG AATCCCAGCA CTCCCACCAG 240 GGACCCCACC CCCGCAGTTT CCAGGATTTC CTCCACCTGT ACCTCCAGGG ACCCCAATGA 300 TTCCTGTACC AATGAGCATT ATGGCTCCTG CTCCGACTGT CTTAGTACCC ACTGTGTCTA 360 TGGTTGGAAA GCATTTGGGC GCAAGAAAGG ATCATCCAGG CTTAAAGGCT AAAGAAAATG 420 ATGAAAATTG TGGTCCTACT ACCACTGTTT TTGTTGGCAA CATTTCCGAG AAAGCTTCAG 480 ACATGCTTAT AAGACAACTC TTAGCTAAAT GTGGTTTGGT TTTGAGCTGG AAGAGAGTAC 540 AAGGTGCTTC CGGAAAGCTT CAAGCCTTCG GATTCTGTGA GTACAAGGAG CCAGAATCTA 600 CCCTCCGTGC ACTCAGATTA TTACATGACC TGCAAATTGG AGAGAAAAAG CTACTCGTTA 660

AAGTTGATGC AAAGACAAAG GCACAGCTGG ATGAATGGAA AGCAAAGAAG AAAGCTTCTA 720 ATGGGAATGC AAGGCCAGAA ACTGTCACTA ATGACGATGA AGAAGCCTTG GATGAAGAAA 780 CAAAGAGGAG AGATCAGATG ATTAAAGGGG CTATTGAAGT TTTAATTCGT GAATACTCCA 840 GTGAGCTAAA TGCCCCCTCA CAGGAATCTG ATTCTCACCC CAGGAAGAAG AAGAAGGAAA 900 AGAAGGAGGA CATTTTCGGC AGATTTCAGT GGGCCCACTG AT 942

#### (2) INFORMATION FOR SEQ ID NO: 153:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2060 base pairs

  - (B) TYPE: nucleic acid(C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (vii) IMMEDIATE SOURCE:
  - (A) LIBRARY: LNODNOT05
  - (B) CLONE: 3125156
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 153:

TCCCCCCTC	AGCCTCCCCC	CCCCCACTG	GCATATGGTC	CTGCCCCTTC	TACCAGACCC	60
ATGGGCCCCC	AGGCAGCCCC	TCTTACCATT	CGAGGGCCCT	CGTCTGCTGG	CCAGTCCACC	120
CCTAGTCCCC	ACCTGGTGCC	TTCACCTGCC	CCATCTCCAG	GGCCTGGTCC	GGTACCCCCT	180
CGCCCCCAG	CAGCAGAACC	ACCCCCTTGC	CTGCGCCGAG	GCGCCGCAGC	TGCAGACCTG	240
CTCTCCTCCA	GCCCGGAGAG	CCAGCATGGC	GGCACTCAGT	CTCCTGGGGG	TGGGCAGCCC	300
CTGCTGCAGC	CCACCAAGGT	GGATGCAGCT	GAGGGTCGTC	GGCCGCAGGC	CCTGCGGCTG	360
ATTGAGCGGG	ACCCCTATGA	GCATCCTGAG	AGGCTGCGGC	AGTTGCAGCA	GGAGCTGGAG	420
GCCTTTCGGG	GTCAGCTGGG	GGATGTGGGA	GCTCTGGACA	CTGTCTGGCG	AGAGCTGCAA	480
GATGCGCAGG	AACATGATGC	CCGAGGCCGT	TCCATCGCCA	TTGCCCGCTG	CTACTCACTG	540
AAGAACCGGC	ACCAGGATGT	CATGCCCTAT	GACAGTAACC	GTGTGGTGCT	GCGCTCAGGC	600
AAGGATGACT	ACATCAATGC	CAGCTGCGTG	GAGGGGCTCT	CCCCATACTG	CCCCCGCTA	660
GTGGCAACCC	AGGCCCCACT	GCCTGGCACA	GCTGCTGACT	TCTGGCTCAT	GGTCCATGAG	720
CAGAAAGTGT	CAGTCATTGT	CATGCTGGTT	TCTGAGGCTG	AGATGGAGAA	GCAAAAAGTG	780
GCACGCTACT	TCCCCACCGA	GAGGGGCCAG	CCCATGGTGC	ACGGTGCCCT	GAGCCTGGCA	840
TTGAGCAGCG	TCCGCAGCAC	CGAAACCCAT	GTGGAGCGCG	TGCTGAGCCT	GCAGTTCCGA	900
GACCAGAGCC	TCAAGCGCTC	TCTTGTGCAC	CTGCACTTCC	CCACTTGGCC	TGAGTTAGGC	960
CTGCCCGACA	GCCCCAGCAA	CTTGCTGCGC	TTCATCCAGG	AGGTGCACGC	ACATTACCTG	1020

CATCAGCGGC CGCTGCACAC GCCCATCATT GTGCACTGCA GCTCTGGTGT GGGCCGCACG 1080 GGAGCCTTTG CACTGCTCTA TGCAGCTGTG CAGGAGGTGG AGGCTGGGAA CGGAATCCCT 1140 GAGCTGCCTC AGCTGGTGCG GCGCATGCGG CAGCAGAGAA AGCACATGCT GCAGGAGAAG 1200 CTGCACCTCA GGTTCTGCTA TGAGGCAGTG GTGAGACACG TGGAGCAGGT CCTGCAGCGC 1260 CATGGTGTGC CTCCTCCATG CAAACCCTTG GCCAGTGCAA GCATCAGCCA GAAGAACCAC 1320 CTTCCTCAGG ACTCCCAGGA CCTGGTCCTC GGTGGGGATG TGCCCATCAG CTCCATCCAG 1380 GCCACCATTG CCAAGCTCAG CATTCGGCCT CCTGGGGGGT TGGAGTCCCC GGTTGCCAGC 1440 TTGCCAGGCC CTGCAGAGCC CCCAGGCCTC CCGCCAGCCA GCCTCCCAGA GTCTACCCCA 1500 ATCCCATCTT CCTCCCAAAC CCCCTTTCCT CCCCACTACC TGAGGCTCCC CAGCCTAAGG 1560 AGGAGCCGCC AGTGCCTGAA GCCCCCAGCT CGGGGCCCCC CTCCTCCTCC CTGGAATTGC 1620 TGGCCTCCTT GACCCCAGAG GCCTTCTCCC TGGACAGCTC CCTGCGGGGC AAACAGCGGA 1680 TGAGCAAGCA TAACTTTCTG CAGGCCCATA ACGGGCAAGG GCTGCGGGCC ACCCGGCCCT 1740 CTGACGACCC CCTCAGCCTT CTGGATCCAC TCTGGACACT CAACAAGACC TGAACAGGTT 1800 TTGCCTACCT GGTCCTTACA CTACATCATC ATCATCTCAT GCCCACCTGC CCACACCCAG 1860 CAGAGCTTCT CAGTGGGCAC AGTCTCTTAC TCCCATTTCT GCTGCCTTTG GCCCTGCCTG 1920 GCCCAGCCTG CACCCCTGTG GGGTGGAAAT GTACTGCAGG CTCTGGGTCA GGTTCTGCTC 1980 CTTTATGGGA CCCGACATTT TTCAGCTCTT TGCTATTGAA ATAATAAACC ACCCTGTTCT 2040 GTGAAAAAA AAAAAAAAG 2060

#### (2) INFORMATION FOR SEO ID NO: 154

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 2065 base pairs
  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

#### (vii) IMMEDIATE SOURCE:

- (A) LIBRARY: LUNGTUT12
- (B) CLONE: 3129120
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 154:

CGGGTCCCCG GGTCTGACAG GAGCAGCCTG TGGGCACCGC GGCGGTAGTT GGAGGCGGGA 60
GAGGGTCCGT AGCCGCCCG CCCTGCCCCG CCATGGGCCT CCTGTCGGAC CCGGTTCGCC 120
GGCGCGCGCT CGCCCGCCTA GTGCTGCCC TCAACGCGCC GTTGTGCGTG CTGAGCTACG 180
TGGCGGGCAT CGCCTGGTTC TTGGCGCTGG TTTTCCCGCC GCTGACCCAG CGCACTTACA 240

TGTCGGAGAA	CGCCATGGGC	TCCACCATGG	TGGAGGAGCA	GTTTGCGGGC	GGAGACCGTG	300
CCCGGGCTTT	TGCCCGGGAC	TTCGCCGCCC	ACCGCAAGAA	GTCGGGGGCT	CTGCCAGTGG	360
CCTGGCTTGA	ACGGACGATG	CGGTCAGTAG	GGCTGGAGGT	CTACACGCAG	AGTTTCTCCC	420
GGAAACTGCC	CTTCCCAGAT	GAGACCCACG	AGCGCTATAT	GGTGTCGGGC	ACCAACGTGT	480
ACGGCATCCT	GCGGGCCCCG	CGTGCTGCCA	GCACCGAGTC	GCTTGTGCTC	ACCGTGCCCT	540
GTGGCTCTGA	CTCTACCAAC	AGCCAGGCTG	TGGGGCTGCT	GCTGGCACTG	GCTGCCCACT	600
TCCGGGGGCA	GATTTATTGG	GCCAAAGATA	TCGTCTTCCT	GGTAACAGAA	CATGACCTTC	660
TGGGCACTGA	GGCTTGGCTT	GAAGCCTACC	ACGATGTCAA	TGTCACTGGC	ATGCAGTCGT	720
CTCCCCTGCA	GGGCCGAGCT	GGGGCCATTC	AGGCAGCCGT	GGCCCTGGAG	CTGAGCAGTG	780
ATGTGGTCAC	CAGCCTCGAT	GTGGCCGTGG	AGGGGCTTAA	CGGGCAGCTG	CCCAACCTTG	840
ACCTGCTCAA	TCTCTTCCAG	ACCTTCTGCC	AGAAAGGGGG	CCTGTTGTGC	ACGCTTCAGG	900
GCAAGCTGCA	GCCCGAGGAC	TGGACATCAT	TGGATGGACC	GCTGCAGGGC	CTGCAGACAC	960
TGCTGCTCAT	GGTTCTGCGG	CAGGCCTCCG	GCCGCCCCA	CGGCTCCCAT	GGCCTCTTCC	1020
TGCGCTACCG	TGTGGAGGCC	CTAACCCTGC	GTGGCATCAA	TAGCTTCCGC	CAGTACAAGT	1080
ATGACCTGGT	GGCAGTGGGC	AAGGCTTTGG	AGGGCATGTT	CCGCAAGCTC	AACCACCTCC	1140
TGGAGCGCCT	GCACCAGTCC	TTCTTCCTCT	ACTTGCTCCC	CGGCCTCTCC	CGCTTCGTCT	1200
CCATCGGCCT	CTACATGCCC	GCTGTCGGCT	TCTTGCTCCT	GGTCCTTGGT	CTCAAGGCTC	1260
TGGAACTGTG	GATGCAGCTG	CATGAGGCTG	GAATGGGCCT	TGAGGAGCCC	GGGGGTGCCC	1320
CTGGCCCCAG	TGTACCCCTT	CCCCCATCAC	AGGGTGTGGG	GCTGGCCTCG	CTCGTGGCAC	1380
CTCTGCTGAT	CTCACAGGCC	ATGGGACTGG	CCCTCTATGT	CCTGCCAGTG	CTGGGCCAAC	1440
ACGTTGCCAC	CCAGCACTTC	CCAGTGGCAG	AGGCTGAGGC	TGTGGTGCTG	ACACTGCTGG	1500
CGATTTATGC	AGCTGGCCTG	GCCCTGCCCC	ACAATACCCA	CCGGGTGGTA	AGCACACAGG	1560
CCCCAGACAG	GGGCTGGATG	GCACTGAAGC	TGGTAGCCCT	GATCTACCTA	GCACTGCAGC	1620
TGGGCTGCAT	CGCCCTCACC	AACTTCTCAC	TGGGCTTCCT	GCTGGCCACC	ACCATGGTGC	1680
CCACTGCTGC	GCTTGCCAAG	CCTCATGGGC	CCCGGACCCT	CTATGCTGCC	CTGCTGGTGC	1740
TGACCAGCCC	GGCAGCCACG	CTCCTTGGCA	GCCTGTTCCT	GTGGCGGGAG	CTGCAGGAGG	1800
CGCCACTGTC	ACTGGCCGAG	GGCTGGCAGC	TCTTCCTGGC	AGCGCTAGCC	CAGGGTGTGC	1860
TGGAGCACCA	CACCTACGGC	GCCCTGCTCT	TCCCACTGCT	GTCCCTGGGC	CTCTACCCCT	1920
GCTGGCTGCT	TTTCTGGAAT	GTGCTCTTCT	GGAAGTGAGA	TCTGCCTGTC	CGGGCTGGGA	1980
CAGAGACTCC	CCAAGGACCC	CATTCTGCCT	CCTTCTGGGG	AAATAAATGA	GTGTCTGTTT	2040
CAGCAGCTAT	TTGATGCTTG	TCACA				2065

<sup>4</sup> 20

25

30

5

10

#### PF-0459 US

### What is claimed is:

- A substantially purified human signal peptide-containing protein (SIGP) 1. comprising a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:1, SEQ ID NO:2, SEQ ID NO:3, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:6, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:10, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, SEQ ID NO:14, SEQ ID NO:15, SEQ ID NO:16, SEQ ID NO:17, SEQ ID NO:18, SEQ ID NO:19, SEQ ID NO:20, SEQ ID NO:21, SEQ ID NO:22, SEQ ID NO:23, SEQ ID NO:24, SEQ ID NO:25, SEQ ID NO:26, SEQ ID NO:27, SEQ ID NO:28, SEQ ID NO:29, SEQ ID NO:30, SEQ ID NO:31, SEQ ID NO:32, SEQ ID NO:33, SEQ ID NO:34, SEO ID NO:35, SEO ID NO:36, SEO ID NO:37, SEO ID NO:38, SEO ID NO:39, SEQ ID NO:40, SEQ ID NO:41, SEQ ID NO:42, SEQ ID NO:43, SEQ ID NO:44, SEQ ID NO:45, SEQ ID NO:46, SEQ ID NO:47, SEQ ID NO:48, SEQ ID NO:49, SEQ ID NO:50, SEQ ID NO:51, SEQ ID NO:52, SEQ ID NO:53, SEQ ID NO:54, SEQ ID NO:55, SEQ ID NO:56, SEQ ID NO:57, SEQ ID NO:58, SEQ ID NO:59, SEQ ID NO:60, SEQ ID NO:61, SEQ ID NO:62, SEQ ID NO:63, SEQ ID NO:64, SEQ ID NO:65, SEQ ID NO:66, SEQ ID NO:67, SEQ ID NO:68, SEQ ID NO:69, SEQ ID NO:70, SEQ ID NO:71, SEQ ID NO:72, SEQ ID NO:73, SEQ ID NO:74, SEQ ID NO:75, SEQ ID NO:76, and SEQ ID NO:77.
- 2. An isolated and purified polynucleotide which hybridizes under stringent conditions to the polynucleotide encoding an SIGP of claim 1.
  - 3. An isolated and purified polynucleotide encoding the SIGP of claim 1.
- 4. A microarray containing at least a fragment of at least one of the polynucleotides encoding an SIGP of claim 1.
- 5. An isolated and purified polynucleotide variant having at least 90% polynucleotide identity to the polynucleotide of claim 3.
  - 6. A composition comprising the polynucleotide of claim 3.
- 7. An isolated and purified polynucleotide which hybridizes under stringent conditions to the polynucleotide of claim 3.
- 8. An isolated and purified polynucleotide which is complementary to the polynucleotide of claim 3.
  - 9. An isolated and purified polynucleotide having a nucleic acid sequence

₹ 20

25

5

#### PF-0459 US

selected from the group consisting of SEQ ID NO:76, SEQ ID NO:77, SEQ ID NO:78, SEQ ID NO:79, SEO ID NO:80, SEQ ID NO:81, SEQ ID NO:82, SEQ ID NO:83, SEQ ID NO:84, SEO ID NO:85, SEO ID NO:86, SEO ID NO:87, SEO ID NO:88, SEO ID NO:89, SEO ID NO:90, SEQ ID NO:91, SEQ ID NO:92, SEQ ID NO:93, SEQ ID NO:94, SEQ ID NO:95, SEQ ID NO:96, SEQ ID NO:97, SEQ ID NO:98, SEQ ID NO:99, SEQ ID NO:100, SEQ ID NO:101, SEQ ID NO:102, SEQ ID NO:103, SEQ ID NO:104, SEQ ID NO:105, SEQ ID NO:106, SEQ ID NO:107, SEQ ID NO:108, SEQ ID NO:109, SEQ ID NO:110, SEQ ID NO:111, SEQ ID NO:112, SEQ ID NO:113, SEQ ID NO:114, SEQ ID NO:115, SEQ ID NO:116, SEQ ID NO:117, SEQ ID NO:118, SEQ ID NO:119, SEQ ID NO:120, SEQ ID NO:121, SEQ ID NO:122, SEQ ID NO:123, SEQ ID NO:124, SEQ ID NO:125, SEQ ID NO:126, SEQ ID NO:127, SEQ ID NO:128, SEQ ID NO:129, SEQ ID NO:130, SEQ ID NO:131, SEQ ID NO:132, SEQ ID NO:133, SEQ ID NO:134, SEQ ID NO:135, SEQ ID NO:136, SEQ ID NO:137, SEQ ID NO:138, SEQ ID NO:139, SEQ ID NO:140, SEQ ID NO:141, SEQ ID NO:142, SEQ ID NO:143, SEQ ID NO:144, SEQ ID NO:145, SEQ ID NO:146, SEQ ID NO:147, SEQ ID NO:148, SEQ ID NO:149, SEQ ID NO:150, SEQ ID NO:151, SEQ ID NO:152, SEQ ID NO:153, and SEQ ID NO:154.

- 10. An isolated and purified polynucleotide variant having at least 90% polynucleotide identity to the polynucleotide of claim 9.
- 11. An isolated and purified polynucleotide which is complementary to the polynucleotide sequence of claim 9.
- 12. An expression vector containing at least a fragment of the polynucleotide of claim 3.
  - 13. A host cell containing the expression vector of claim 12.
- 14. A method for producing a polypeptide encoding a human signal peptidecontaining protein, the method comprising the steps of:
  - (a) culturing the host cell of claim 13 under conditions suitable for the expression of the polypeptide; and
    - (b) recovering the polypeptide from the host cell culture.
- 15. A pharmaceutical composition comprising the SIGP of claim 1 in conjunction with a suitable pharmaceutical carrier.

5

10

- 16. A purified antibody which specifically binds to the SIGP of claim 1.
- 17. A purified agonist of the SIGP of claim 1.
- 18. A purified antagonist of the SIGP of claim 1.
- 19. A method for treating or preventing a cancer, the method comprising administering to a subject in need of such treatment an effective amount of the pharmaceutical composition of claim 15.
- 20. A method for treating or preventing a cancer, the method comprising administering to a subject in need of such treatment an effective amount of the antagonist of claim 18.
- 21. A method for treating or preventing an immune response, the method comprising administering to a subject in need of such treatment an effective amount of the antagonist of claim 18.
- 22. A method for detecting a polynucleotide encoding a human signal peptidecontaining protein in a biological sample containing nucleic acids, the method comprising the steps of:
  - (a) hybridizing the polynucleotide of claim 8 to at least one of the nucleic acids of the biological sample, thereby forming a hybridization complex; and
  - (b) detecting the hybridization complex, wherein the presence of the hybridization complex correlates with the presence of a polynucleotide encoding SIGP in the biological sample.
- 23. The method of claim 22 wherein the nucleic acids of the biological sample are amplified by the polymerase chain reaction prior to the hybridizing step.

## ABSTRACT OF THE DISCLOSURE

The invention provides a human signal peptide-containing proteins (SIGP) and polynucleotides which identify and encode SIGP. The invention also provides expression vectors, host cells, antibodies, agonists, and antagonists. The invention also provides methods for treating or preventing disorders associated with expression of SIGP.



# DECLARATION AND POWER OF ATTORNEY FOR UNITED STATES PATENT APPLICATION

As a below named inventor, I hereby declare that:

My residence, post office address and citizenship are as stated below next to my name, and

I believe that I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if more than one name is listed below) of the subject matter which is claimed and for which a United States patent is sought on the invention entitled

## **HUMAN SIGNAL PEPTIDE-CONTAINING PROTEINS**

the specification of which:	
/_/ is attached hereto.	
/X/ was filed on December 31, 1997, as application Serial No. 09/002,485 and if this box contains an X //, was amended on	
/_/ was filed as Patent Cooperation Treaty international application No on, 19, if this box contains an X /_/, was amended on under Patent Cooperation Treaty Article 19 on19, and if this box contains an X /_/, was amended on	-
I hereby state that I have reviewed and understand the contents of the above-identified specification, including the claims, as amended by any amendment referred to above.	
I acknowledge my duty to disclose information which is material to the examination of	

I hereby claim the benefit under Title 35, United States Code, §119 or §365(a)-(b) of any foreign application(s) for patent or inventor's certificate indicated below and of any Patent Cooperation Treaty international applications(s) designating at least one country other than the United States indicated below and have also identified below any foreign application(s) for patent or inventor's certificate and Patent Cooperation Treaty international application(s) designating at least one country other than the United States for the same subject matter and having a filing date before that of the application for said subject matter the priority of which is claimed:

this application in accordance with Title 37, Code of Federal Regulations, §1.56(a).

Country	Number	Filing Date	Priority Claimed
			/_/ Yes /_/ No
			/_/ Yes /_/ No

I hereby claim the benefit under Title 35, United States Code, §119(e) of any United States provisional application(s) listed below.

I hereby claim the benefit under Title 35, United States Code, §120 of any United States application(s) listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in said prior application(s) in the manner required by the first paragraph of Title 35, United States Code §112, I acknowledge my duty to disclose material information as defined in Title 37 Code of Federal Regulations, §1.56(a) which occurred between the filing date(s) of the prior application(s) and the national or Patent Cooperation Treaty international filing date of this application:

Application		Status (Pending,
Serial No.	Filed	Abandoned, Patented)

I hereby appoint the following:

LUCY J. BILLINGS
MICHAEL C. CERRONE
SHEELA MOHAN-PETERSON
COLETTE C. MUENZEN
KAREN J. ZELLER

Registration No. 36,749 Registration No. 39,132 Registration No. 41,201 Registration No. 39,784 Registration No. 37,071

respectively and individually, as my attorneys and/or agents, with full power of substitution and revocation, to prosecute this application and to transact all business in the Patent and Trademark Office connected therewith. Please address all communications to:

MICHAEL C. CERRONE, Ph.D. INCYTE PHARMACEUTICALS, INC. 3174 PORTER DRIVE, PALO ALTO, CA 94304

TEL: 650-855-0555 FAX: 650-845-4166

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are

punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issuing thereon.

**Sole Inventor or First Joint Inventor:** 

Full name:

PREETI LAL

Signature:

Presti Cel 15th May 1998

Date:

India

Residence:

Citizenship:

Santa Clara, California

P.O. Address:

2382 Lass Drive

Santa Clara, CA 95054

**Scond Joint Inventor:** 

Full name:

JENNIFER L. HILLMAN

Signature:

Date:

United States of America

Citizenship: Residence:

Mountain View, California

P.O. Address:

230 Monroe Drive, #12

Mountain View, CA 94040

The state of the s

<b>Third Joint Inventor:</b> Full name:
---

NEIL C. CORLE

Signature:

Date:

United States of America

Residence:

Citizenship:

Mountain View, California

P.O. Address:

1240 Dale Avenue, #30

Mountain View, CA 94040

**Fourth Joint Inventor:** 

Full name:

KARL J. GUEGLER

Signature:

Date:

Switzerland

Residence:

Citizenship:

Menlo Park, California

P.O. Address:

1048 Oakland Avenue

Menlo Park, CA 94025

**Fifth Joint Inventor:** 

Full Name:

BAUGHN MARIAH R. <del>BAUGH</del>

Signature:

Date:

Citizenship:

United States of America

Residence:

San Jose, California

P.,O. Address:

1537 LaRossa Circle

San Jose, CA 95125

Sixth Joint Inventor: Full Name: SUSAN K. SATHER

Signature: Susan K. Nather

Date: 5/18/98

Citizenship: United States of America

Residence: Palo Alto, California

P.O. Address: 2721 Midtown Court, Apt. #203

Palo Alto, CA 94303

Seventh Joint Inventor: Full name: PURVI SHAH

Signature: "PuniShah

Date: \_\_\_\_\_\_5/15/98\_\_\_\_\_

Citizenship: India

Residence: Sunnyvale, California

P.O. Address: 1608 Queen Charlotte Drive, #5

Sunnyvale, CA 94087